

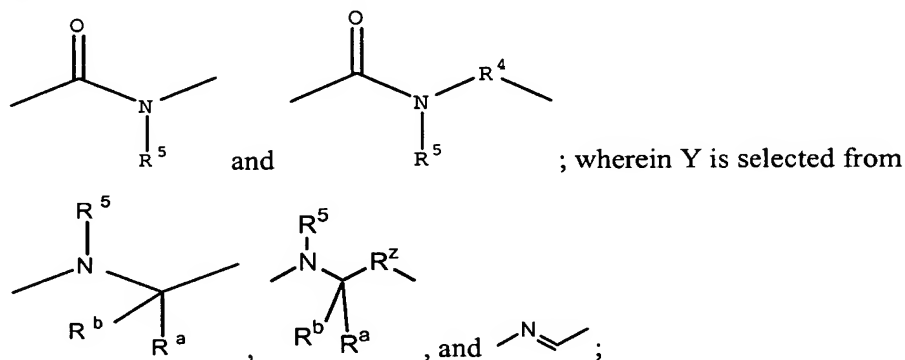
The listing of claims will replace all prior versions, and listings, of claims in the application:

### Listing of Claims

Claim 1 (canceled).

Claim 2 (withdrawn): Compound of Claim 1, and pharmaceutically acceptable salts thereof, wherein A is selected from 5- or 6- membered partially saturated heterocyclyl.

Claim 3 (withdrawn): Compound of Claim 2, and pharmaceutically acceptable salts thereof, wherein A is selected from dihydropyran, dihydrothienyl, dihydrofuryl, oxo-dihydrofuryl, pyrrolinyl, dihydrothiazolyl, dihydro-oxazolyl, dihydro-isothiazolyl, dihydro-isoxazolyl, imidazoliny and pyrazoliny; wherein X is selected from



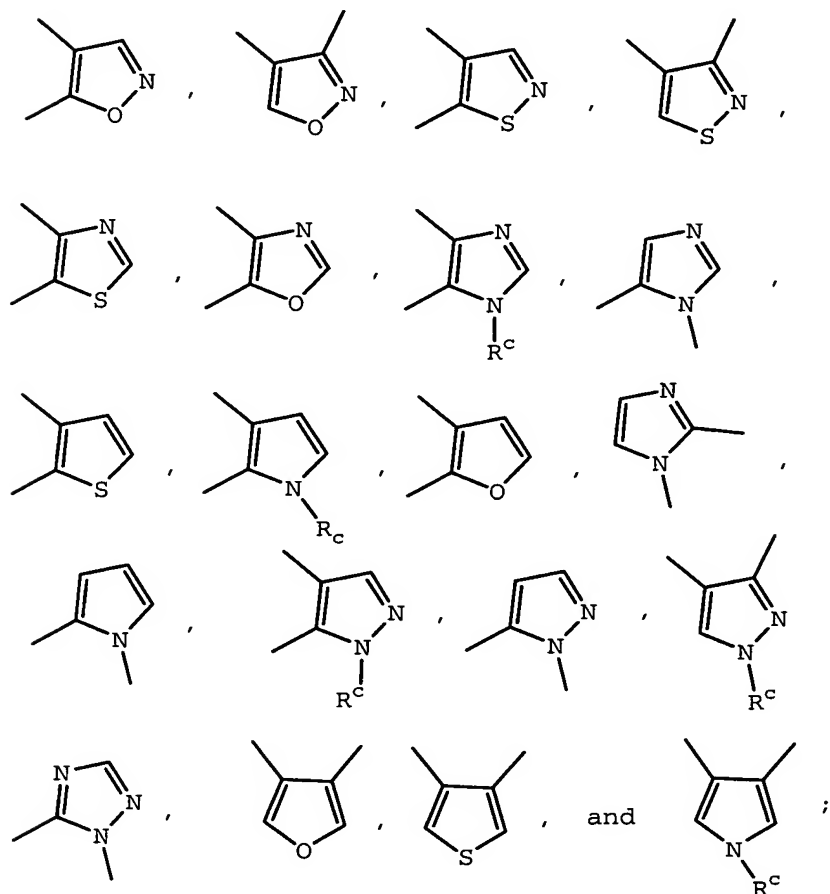
wherein  $\text{R}^a$  and  $\text{R}^b$  are independently selected from H, halo, cyano, and  $\text{C}_{1-2}$ -alkyl substituted with  $\text{R}^2$ , or wherein  $\text{R}^a$  and  $\text{R}^b$  together form  $\text{C}_3\text{-C}_4$  cycloalkyl; wherein  $\text{R}^z$  is  $\text{C}_1\text{-C}_2$  alkylene, where one of the  $\text{CH}_2$  groups may be substituted with an oxygen atom or an  $-\text{NH}-$ ; wherein R is selected from substituted or unsubstituted 5-6 membered heteroaryl comprising one or more nitrogen atoms, and substituted or unsubstituted 9-10 membered fused heteroaryl comprising one or more nitrogen atoms; wherein substituted R is substituted with one or more substituents independently selected from halo,  $-\text{OR}^3$ ,  $-\text{SR}^3$ ,  $-\text{CO}_2\text{R}^3$ ,  $-\text{CONR}^3\text{R}^3$ ,  $-\text{COR}^3$ ,  $-\text{NR}^3\text{R}^3$ ,  $-\text{SO}_2\text{NR}^3\text{R}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{OR}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{R}^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $\text{C}_{1-2}$ -alkyl, cyano,  $\text{C}_{1-2}$ -hydroxyalkyl, nitro and  $\text{C}_{1-2}$ -haloalkyl; wherein  $\text{R}^1$  is selected from substituted or unsubstituted aryl selected from phenyl, naphthyl, indenyl and tetrahydronaphthyl, substituted or unsubstituted 5-6 membered heteroaryl, and substituted or unsubstituted 9-10 membered fused heteroaryl; wherein substituted  $\text{R}^1$  is substituted with one or more substituents independently selected from halo,  $-\text{OR}^3$ ,  $-\text{SR}^3$ ,  $-\text{SO}_2\text{R}^3$ ,  $-\text{CO}_2\text{R}^3$ ,  $-\text{CONR}^3\text{R}^3$ ,  $-\text{COR}^3$ ,  $-\text{NR}^3\text{R}^3$ ,  $-\text{NH}(\text{C}_1\text{-C}_2 \text{ alkyleneR}^3)$ ,  $-(\text{C}_1\text{-C}_2 \text{ alkylene})\text{NR}^3\text{R}^3$ ,  $-\text{SO}_2\text{NR}^3\text{R}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{OR}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{R}^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, optionally substituted phenyl- $\text{C}_{1-2}$ -alkylene, optionally substituted 5-6 membered heterocyclyl- $\text{C}_{1-2}$ -alkylene,  $\text{C}_{1-2}$ -alkyl, cyano,  $\text{C}_{1-2}$ -hydroxyalkyl, nitro and  $\text{C}_{1-2}$ -haloalkyl; wherein  $\text{R}^2$  is one or more substituents independently selected

from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl,  $C_{1-3}$ -carboxyalkyl, nitro,  $C_{2-3}$ -alkenyl,  $C_{2-3}$ -alkynyl and  $C_{1-2}$ -haloalkyl; wherein  $R^3$  is selected from H,  $C_{1-2}$ -alkyl, phenyl,  $C_3-C_6$  cycloalkyl and  $C_{1-2}$ -haloalkyl; wherein  $R^4$  is  $C_{2-3}$ -alkylenyl, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ; and wherein  $R^5$  is selected from H and  $C_{1-2}$ -alkyl.

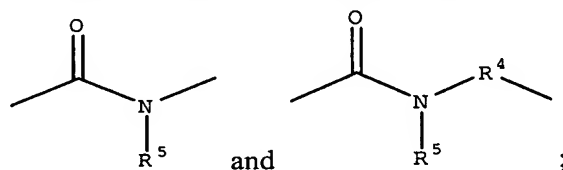
Claim 4 (canceled).

Claim 5 (canceled).

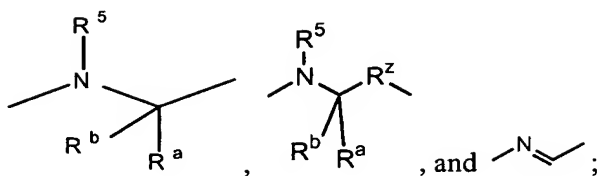
Claim 6 (withdrawn): Compound of Claim 1 wherein A is selected from



wherein  $R^c$  is selected from H, methyl and optionally substituted phenyl; wherein X is selected from

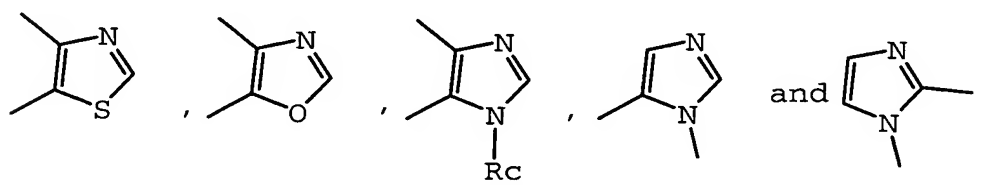


wherein Y is selected from

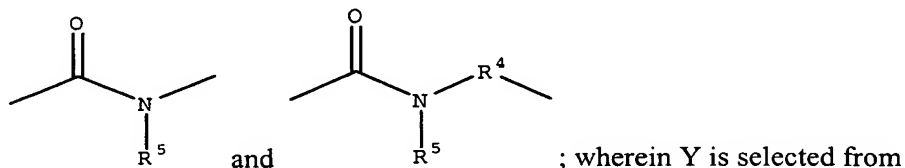


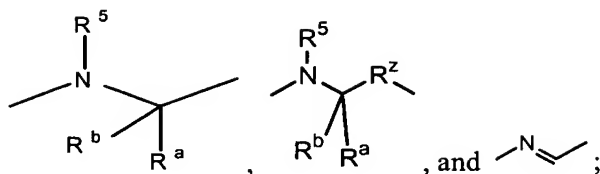
wherein  $R^a$  and  $R^b$  are independently selected from H, halo, cyano, and  $C_{1-2}$ -alkyl substituted with  $R^2$ , or wherein  $R^a$  and  $R^b$  together form  $C_3-C_4$  cycloalkyl; wherein  $R^z$  is  $C_1-C_2$  alkylene, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ; wherein  $R$  is selected from substituted or unsubstituted 5-6 membered heteroaryl comprising one or more nitrogen atoms, and substituted or unsubstituted 9-10 membered fused heteroaryl comprising one or more nitrogen atoms; wherein substituted  $R$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^1$  is selected from substituted or unsubstituted aryl selected from phenyl, naphthyl, indenyl and tetrahydronaphthyl, substituted or unsubstituted 5-6 membered heteroaryl, and substituted or unsubstituted 9-10 membered fused heteroaryl; wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-SO_2R^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-NH(C_{1-2} \text{ alkylene} R^3)$ ,  $-(C_{1-2} \text{ alkylene})NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-2}$ -alkylene, optionally substituted 5-6 membered heterocyclyl- $C_{1-2}$ -alkylene,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl,  $C_{1-3}$ -carboxyalkyl, nitro,  $C_{2-3}$ -alkenyl,  $C_{2-3}$ -alkynyl and  $C_{1-2}$ -haloalkyl; wherein  $R^3$  is selected from H,  $C_{1-2}$ -alkyl, phenyl,  $C_3-C_6$  cycloalkyl and  $C_{1-2}$ -haloalkyl; wherein  $R^4$  is  $C_{2-3}$ -alkylene, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ; and wherein  $R^5$  is selected from H and  $C_{1-2}$ -alkyl.

Claim 7 (withdrawn): Compound of Claim 6 wherein A is selected from



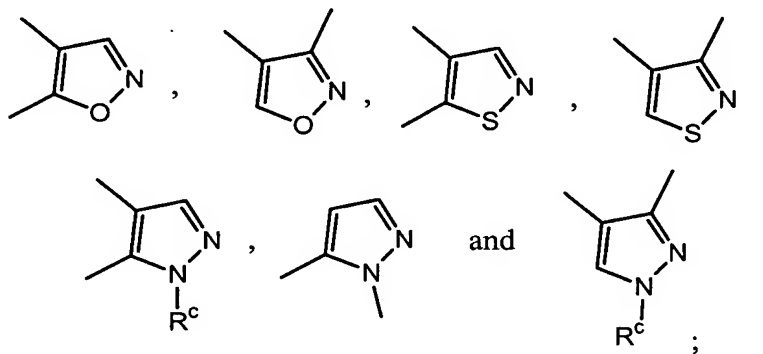
wherein  $R^c$  is selected from H, methyl and optionally substituted phenyl; wherein X is selected from



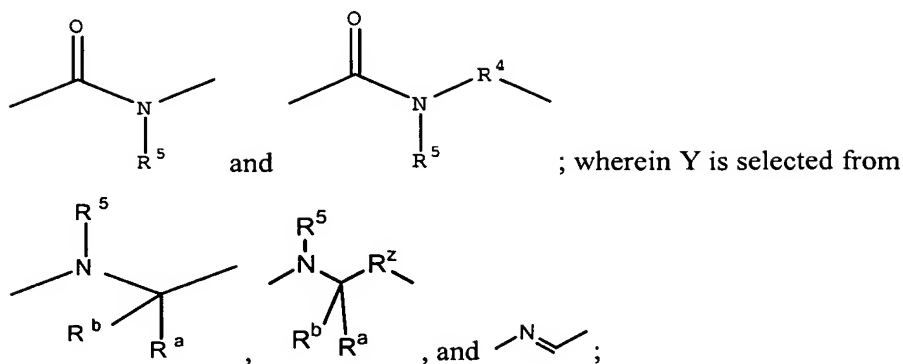


wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, and C<sub>1-2</sub>-alkyl; wherein R<sup>z</sup> is C<sub>1-2</sub> alkylenyl; wherein R is selected from substituted or unsubstituted 4-pyridyl, 4-pyrimidinyl, 5-pyrimidinyl, 6-pyrimidinyl, 4-pyridazinyl, 6-pyridazinyl, indazolyl, quinolinyl, isoquinolinyl, quinazolinyl, triazolyl and 4-pyrazolyl; wherein substituted R is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, C<sub>1-2</sub>-alkyl, cyano, C<sub>1-2</sub>-hydroxyalkyl, nitro and C<sub>1-2</sub>-haloalkyl; wherein R<sup>1</sup> is a substituted or unsubstituted substituent selected from phenyl, indenyl, thienyl, indolyl, pyridyl, naphthyl, tetrahydronaphthyl, 2,1,3-benzothiadiazolyl, indazolyl, quinolyl, isoquinolyl, pyrimidinyl, pyridazinyl, pyrazolyl, imidazolyl, oxazolyl, thiazolyl, thiadiazolyl, tetrahydroquinolinyl, benzodioxanyl, quinazolinyl, furyl and pyrrolyl; wherein substituted R<sup>1</sup> is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -NH(C<sub>1-2</sub>-alkylenyl-R<sup>3</sup>), -(C<sub>1-2</sub>-alkylenyl)NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, optionally substituted phenyl-C<sub>1-2</sub>-alkylenyl, optionally substituted 5-6 membered heterocyclyl-C<sub>1-2</sub>-alkylenyl, C<sub>1-2</sub>-alkyl, cyano, C<sub>1-2</sub>-hydroxyalkyl, nitro and C<sub>1-2</sub>-haloalkyl; wherein R<sup>2</sup> is one or more substituents independently selected from H, halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, C<sub>1-2</sub>-alkyl, cyano, C<sub>1-2</sub>-hydroxyalkyl, C<sub>1-3</sub>-carboxyalkyl, nitro, C<sub>2-3</sub>-alkenyl, C<sub>2-3</sub>-alkynyl and C<sub>1-2</sub>-haloalkyl; wherein R<sup>3</sup> is selected from H, methyl, ethyl, cyclopropyl, cyclohexyl and trifluoromethyl; wherein R<sup>4</sup> is C<sub>2-3</sub>-alkylenyl; and wherein R<sup>5</sup> is from H, methyl or ethyl; and pharmaceutically acceptable salts thereof.

Claim 8 (withdrawn): Compound of Claim 6 wherein A is selected from

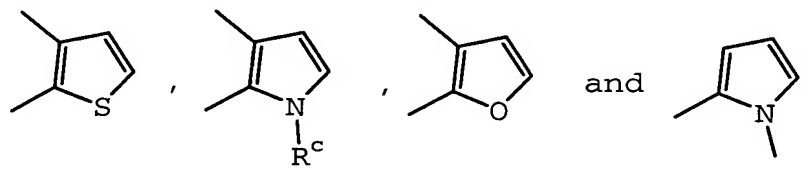


wherein R<sup>c</sup> is selected from H, methyl and optionally substituted phenyl; wherein X is selected from

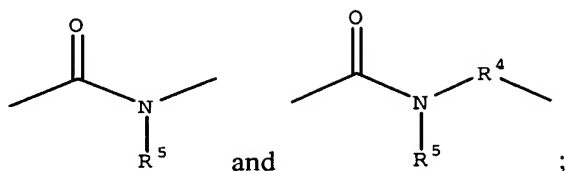


wherein  $R^a$  and  $R^b$  are independently selected from H, halo, and  $C_{1-2}$ -alkyl; wherein  $R^Z$  is  $C_1$ - $C_2$  alkylene; wherein R is selected from substituted or unsubstituted 4-pyridyl, 4-pyrimidinyl, 5-pyrimidinyl, 6-pyrimidinyl, 4-pyridazinyl, 6-pyridazinyl, indazolyl, quinolinyl, isoquinolinyl, quinoxalinyl, triazolyl and 4-pyrazolyl; wherein substituted R is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^1$  is a substituted or unsubstituted substituent selected from phenyl, indenyl, thienyl, indolyl, pyridyl, naphthyl, tetrahydronaphthyl, 2,1,3-benzothiadiazolyl, indazolyl, quinolyl, isoquinolyl, pyrimidinyl, pyridazinyl, pyrazolyl, imidazolyl, oxazolyl, thiazolyl, thiadiazolyl, tetrahydroquinolyl, benzodioxanyl, quinazolinyl, furyl and pyrrolyl; wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-NH(C_{1-2}\text{-alkylene}-R^3)$ ,  $-(C_{1-2}\text{-alkylene})NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-2}$ -alkylene, optionally substituted 5-6 membered heterocyclyl- $C_{1-2}$ -alkylene,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl,  $C_{1-3}$ -carboxyalkyl, nitro,  $C_{2-3}$ -alkenyl,  $C_{2-3}$ -alkynyl and  $C_{1-2}$ -haloalkyl; wherein  $R^3$  is selected from H, methyl, ethyl, cyclopropyl, cyclohexyl and trifluoromethyl; wherein  $R^4$  is  $C_{2-3}$ -alkylene; and wherein  $R^5$  is from H, methyl or ethyl; and pharmaceutically acceptable salts thereof.

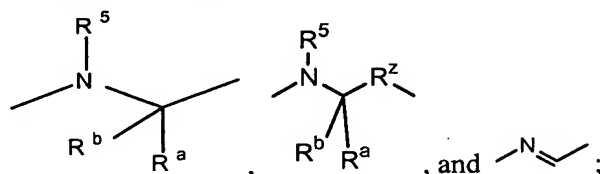
Claim 9 (withdrawn): Compound of Claim 6 wherein A is selected from



wherein  $R^c$  is selected from H, methyl and optionally substituted phenyl; wherein X is selected from



wherein Y is selected from

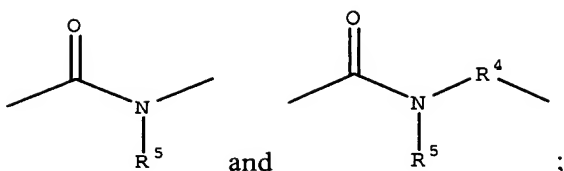


wherein  $R^a$  and  $R^b$  are independently selected from H, halo, and  $C_{1-2}$ -alkyl; wherein  $R^z$  is  $C_1$ - $C_2$  alkylenyl; wherein R is selected from substituted or unsubstituted 4-pyridyl, 4-pyrimidinyl, 5-pyrimidinyl, 6-pyrimidinyl, 4-pyridazinyl, 6-pyridazinyl, indazolyl, quinolinyl, isoquinolinyl, quinazolinyl, triazolyl and 4-pyrazolyl; wherein substituted R is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^1$  is a substituted or unsubstituted substituent selected from phenyl, indenyl, thienyl, indolyl, pyridyl, naphthyl, tetrahydronaphthyl, 2,1,3-benzothiadiazolyl, indazolyl, quinolyl, isoquinolyl, pyrimidinyl, pyridazinyl, pyrazolyl, imidazolyl, oxazolyl, thiazolyl, thiadiazolyl, tetrahydroquinolinyl, benzodioxanyl, quinazolinyl, furyl and pyrrolyl; wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-NH(C_1-C_2\text{-alkylenyl}-R^3)$ ,  $-(C_1-C_2\text{-alkylenyl})NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-2}$ -alkylenyl, optionally substituted 5-6 membered heterocyclyl- $C_{1-2}$ -alkylenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl,  $C_{1-3}$ -carboxyalkyl, nitro,  $C_{2-3}$ -alkenyl,  $C_{2-3}$ -alkynyl and  $C_{1-2}$ -haloalkyl; wherein  $R^3$  is selected from H, methyl, ethyl, cyclopropyl, cyclohexyl and trifluoromethyl; wherein  $R^4$  is  $C_{2-3}$ -alkylenyl; and wherein  $R^5$  is from H, methyl or ethyl; and pharmaceutically acceptable salts thereof.

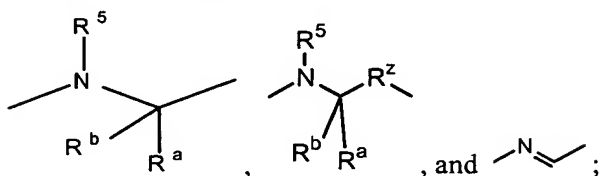
Claim 10 (canceled).

Claim 11 (canceled).

Claim 12 (withdrawn): Compound of Claim 1 wherein A is 9- or 10-membered fused partially saturated heterocyclyl or 9- or 10-membered fused heteroaryl; wherein X is selected from



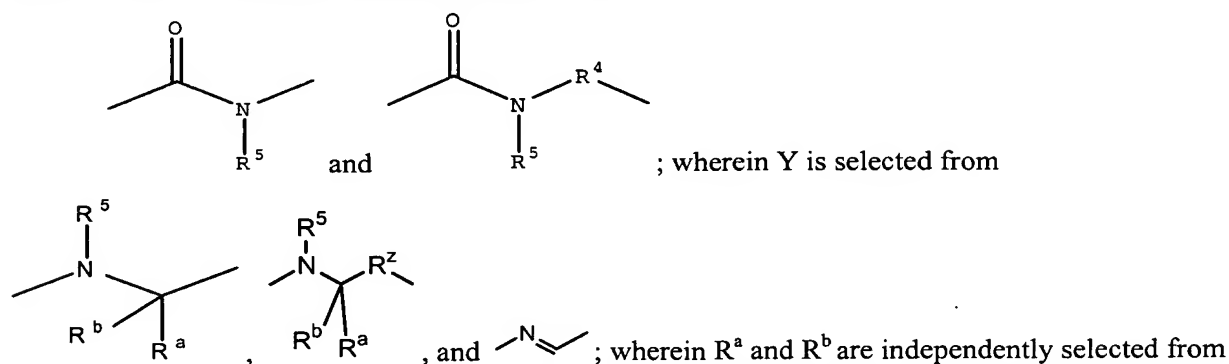
wherein Y is selected from



wherein  $R^a$  and  $R^b$  are independently selected from H, halo, cyano, and  $C_{1-2}$ -alkyl substituted with  $R^2$ , or wherein  $R^a$  and  $R^b$  together form  $C_3$ - $C_4$  cycloalkyl; wherein  $R^z$  is  $C_1$ - $C_2$  alkylene, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an -NH-; wherein R is selected from substituted or unsubstituted 5-6 membered heteroaryl comprising one or more nitrogen atoms, and substituted or unsubstituted 9-10 membered fused heteroaryl comprising one or more nitrogen atoms; wherein substituted R is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, cycloalkyl, optionally substituted 5-6 membered heterocycl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^1$  is selected from substituted or unsubstituted aryl selected from phenyl, naphthyl, indenyl and tetrahydronaphthyl, substituted or unsubstituted 5-6 membered heteroaryl, and substituted or unsubstituted 9-10 membered fused heteroaryl; wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -NH( $C_{1-2}$  alkyleneR<sup>3</sup>), -( $C_{1-2}$  alkylene)NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocycl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-2}$ -alkylene, optionally substituted 5-6 membered heterocycl- $C_{1-2}$ -alkylene,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^2$  is one or more substituents independently selected from H, halo, -OR<sup>3</sup>, oxo, -SR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, cycloalkyl, optionally substituted 5-6 membered heterocycl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl,  $C_{1-3}$ -carboxyalkyl, nitro,  $C_{2-3}$ -alkenyl,  $C_{2-3}$ -alkynyl and  $C_{1-2}$ -haloalkyl; wherein  $R^3$  is selected from H,  $C_{1-2}$ -alkyl, phenyl,  $C_3$ - $C_6$  cycloalkyl and  $C_{1-2}$ -haloalkyl; wherein  $R^4$  is  $C_{2-3}$ -alkylene, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an -NH-; and wherein  $R^5$  is selected from H and  $C_{1-2}$ -alkyl; and pharmaceutically acceptable salts thereof.

Claim 13 (withdrawn): Compound of Claim 12 wherein A is selected from benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, benzthiazolyl, indazolyl, indolyl, isoindolyl, quinolyl, isoquinolyl, naphthpyridinyl, tetrahydroquinolyl, quinoxalinyl and quinazolinyl; and pharmaceutically acceptable salts thereof.

Claim 14 (withdrawn): Compound of Claim 1, and pharmaceutically acceptable salts thereof, wherein A is 5- or 6-membered cycloalkenyl; wherein X is selected from



, and  $\text{CH}_2=\text{CH}-$ ; wherein  $R^a$  and  $R^b$  are independently selected from H, halo, cyano, and  $C_{1-2}$ -alkyl substituted with  $R^2$ , or wherein  $R^a$  and  $R^b$  together form  $C_3-C_4$  cycloalkyl; wherein  $R^Z$  is  $C_1-C_2$  alkylene, where one of the  $\text{CH}_2$  groups may be substituted with an oxygen atom or an  $-\text{NH}-$ ; wherein R is selected from substituted or unsubstituted 5-6 membered heteroaryl comprising one or more nitrogen atoms, and substituted or unsubstituted 9-10 membered fused heteroaryl comprising one or more nitrogen atoms; wherein substituted R is substituted with one or more substituents independently selected from halo,  $-\text{OR}^3$ ,  $-\text{SR}^3$ ,  $-\text{CO}_2\text{R}^3$ ,  $-\text{CONR}^3\text{R}^3$ ,  $-\text{COR}^3$ ,  $-\text{NR}^3\text{R}^3$ ,  $-\text{SO}_2\text{NR}^3\text{R}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{OR}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{R}^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^1$  is selected from substituted or unsubstituted aryl selected from phenyl, naphthyl, indenyl and tetrahydronaphthyl, substituted or unsubstituted 5-6 membered heteroaryl, and substituted or unsubstituted 9-10 membered fused heteroaryl; wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo,  $-\text{OR}^3$ ,  $-\text{SR}^3$ ,  $-\text{SO}_2\text{R}^3$ ,  $-\text{CO}_2\text{R}^3$ ,  $-\text{CONR}^3\text{R}^3$ ,  $-\text{COR}^3$ ,  $-\text{NR}^3\text{R}^3$ ,  $-\text{NH}(\text{C}_{1-2} \text{ alkylene} \text{R}^3)$ ,  $-(\text{C}_{1-2} \text{ alkylene})\text{NR}^3\text{R}^3$ ,  $-\text{SO}_2\text{NR}^3\text{R}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{OR}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{R}^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-2}$ -alkylene, optionally substituted 5-6 membered heterocyclyl- $C_{1-2}$ -alkylene,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl, nitro and  $C_{1-2}$ -haloalkyl; wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-\text{OR}^3$ , oxo,  $-\text{SR}^3$ ,  $-\text{CO}_2\text{R}^3$ ,  $-\text{CONR}^3\text{R}^3$ ,  $-\text{COR}^3$ ,  $-\text{NR}^3\text{R}^3$ ,  $-\text{SO}_2\text{NR}^3\text{R}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{OR}^3$ ,  $-\text{NR}^3\text{C}(\text{O})\text{R}^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl,  $C_{1-2}$ -alkyl, cyano,  $C_{1-2}$ -hydroxyalkyl,  $C_{1-3}$ -carboxyalkyl, nitro,  $C_{2-3}$ -alkenyl,  $C_{2-3}$ -alkynyl and  $C_{1-2}$ -haloalkyl; wherein  $R^3$  is selected from H,  $C_{1-2}$ -alkyl, phenyl,  $C_3-C_6$  cycloalkyl and  $C_{1-2}$ -haloalkyl; wherein  $R^4$  is  $C_{2-3}$ -alkylene, where one of the  $\text{CH}_2$  groups may be substituted with an oxygen atom or an  $-\text{NH}-$ ; and wherein  $R^5$  is selected from H and  $C_{1-2}$ -alkyl.

Claim 15 (withdrawn): Compound of Claim 14 wherein A is cyclopentadienyl or cyclopentenyl; and pharmaceutically acceptable salts thereof.



Claim 16 (canceled).

Claim 17 (canceled).

Claim 18 (canceled).

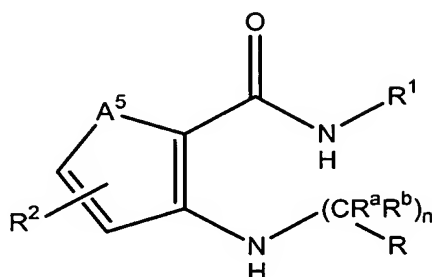
Claim 19 (canceled).

Claim 20 (canceled).

Claim 21 (canceled).

Claim 22 (canceled).

Claim 23 (withdrawn): A compound of Claim 1 having the Formula V



V

wherein  $A^5$  is selected from S, O and  $NR^6$ ;

wherein  $R^a$  and  $R^b$  are independently selected from H, halo,  $C_{1-4}$ -alkyl and  $-N(R^6)_2$ ;

wherein n is 1-2;

wherein R is selected from

a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and

b) unsubstituted or substituted 9- or 10-membered fused nitrogen-containing heteroaryl,

where R is substituted with one or more substituents selected from halo, amino, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl and  $C_{1-6}$ -alkoxy;

wherein  $R^1$  is selected from unsubstituted or substituted

aryl,

5-6 membered heteroaryl and

9-10 membered fused heteroaryl,

wherein substituted  $R^1$  is substituted with one or more substituents selected from halo,  $C_{1-6}$ -alkyl,

optionally substituted  $C_{3-6}$ -cycloalkyl, optionally substituted phenyl,  $C_{1-6}$ -haloalkoxy,

optionally substituted phenyloxy, benzyl, optionally substituted 5-6 membered heterocycl-

C<sub>1</sub>-C<sub>2</sub>-alkylenyl, optionally substituted heteroaryl, optionally substituted heteroaryloxy, C<sub>1-6</sub>-haloalkyl, and C<sub>1-6</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,  
halo,  
C<sub>1-6</sub>-alkyl,  
C<sub>1-6</sub>-haloalkyl,  
C<sub>1-6</sub>-alkoxy,  
C<sub>1-6</sub>-haloalkoxy,  
C<sub>1-6</sub>-carboxyalkyl,  
unsubstituted or substituted aryl and  
unsubstituted or substituted 5-6 membered heteroaryl; and

wherein R<sup>6</sup> is H or C<sub>1-6</sub>-alkyl;

and pharmaceutically acceptable isomers and salts thereof.

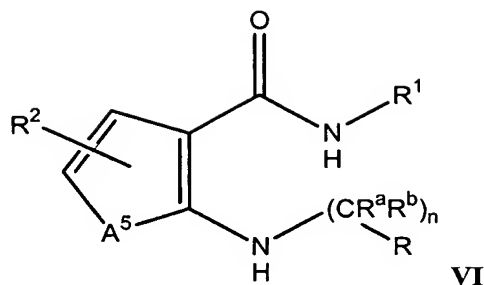
Claim 24 (withdrawn): Compound of Claim 23 wherein R<sup>a</sup> and R<sup>b</sup> are H;  
wherein n is 1-2;

wherein R is selected from 4-pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, indazolyl, quinolyl, isoquinolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy and ethoxy;

wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, naphthyl, isoquinolyl, quinolyl, pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, naphthyridinyl, quinozaliny, tetrahydroquinolyl, indazolyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, or benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, cyclohexyl, phenylmethyl, morpholinylmethyl, methylpiperdinylmethyl, methylpiperazinylmethyl, ethyl, propyl, trifluoromethyl, phenyloxy, methoxy and ethoxy; and

wherein R<sup>2</sup> is one or more substituents independently selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl;  
and pharmaceutically acceptable salts thereof.

Claim 25 (withdrawn): A compound of Claim 1 having the Formula



wherein A<sup>5</sup> is selected from S, O and NR<sup>6</sup>;

wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, C<sub>1-4</sub>-alkyl and -N(R<sup>6</sup>)<sub>2</sub>;

wherein n is 1-2;

wherein R is selected from

a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and

b) unsubstituted or substituted 9- or 10-membered fused nitrogen-containing heteroaryl,

where R is substituted with one or more substituents selected from halo, amino, hydroxy, C<sub>1-6</sub>-alkyl, C<sub>1-6</sub>-haloalkyl and C<sub>1-6</sub>-alkoxy;

wherein R<sup>1</sup> is selected from unsubstituted or substituted

aryl,

5-6 membered heteroaryl and

9-10 membered fused heteroaryl,

wherein substituted R<sup>1</sup> is substituted with one or more substituents selected from halo, C<sub>1-6</sub>-alkyl,

optionally substituted C<sub>3-6</sub>-cycloalkyl, optionally substituted phenyl, C<sub>1-6</sub>-haloalkoxy,

optionally substituted phenyloxy, benzyl, optionally substituted 5-6 membered heterocyclyl-

C<sub>1</sub>-C<sub>2</sub>-alkylenyl, optionally substituted heteroaryl, optionally substituted heteroaryloxy, C<sub>1-6</sub>-haloalkyl, and C<sub>1-6</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,

halo,

C<sub>1-6</sub>-alkyl,

C<sub>1-6</sub>-haloalkyl,

C<sub>1-6</sub>-alkoxy,

C<sub>1-6</sub>-haloalkoxy,

C<sub>1-6</sub>-carboxyalkyl,

unsubstituted or substituted aryl and

unsubstituted or substituted 5-6 membered heteroaryl; and

wherein R<sup>6</sup> is H or C<sub>1-6</sub>-alkyl;

and pharmaceutically acceptable isomers and salts thereof.

Claim 26 (withdrawn): Compound of Claim 25 wherein R<sup>a</sup> and R<sup>b</sup> are H;

wherein n is 1-2;

wherein R is selected from 4-pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, indazolyl, quinolyl, isoquinolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy and ethoxy;

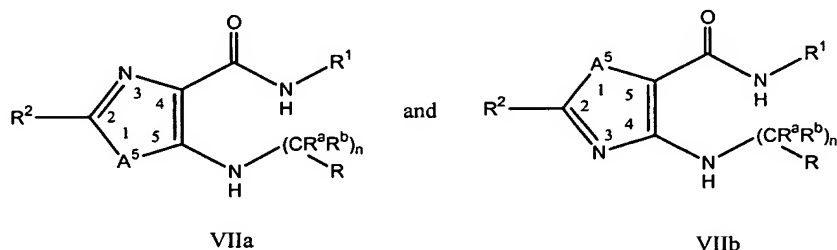
wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, naphthyl, isoquinolyl, quinolyl, pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, naphthyridinyl, quinozaliny, tetrahydroquinolyl, indazolyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, or benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, cyclohexyl, phenylmethyl, morpholinylmethyl, methylpiperdinylmethyl, methylpiperazinylmethyl, ethyl, propyl, trifluoromethyl, phenyloxy, methoxy and ethoxy; and

wherein R<sup>2</sup> is one or more substituents independently selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected

from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl;

and pharmaceutically acceptable salts thereof.

Claim 27 (withdrawn): A compound of Claim 1 having the Formula



wherein A<sup>5</sup> is selected from S, O and NR<sup>6</sup>;

wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, C<sub>1-4</sub>-alkyl and -N(R<sup>6</sup>)<sub>2</sub>;

wherein n is 1-2;

wherein R is selected from

- a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and
  - b) unsubstituted or substituted 9- or 10-membered fused nitrogen-containing heteroaryl,
- where R is substituted with one or more substituents selected from halo, amino, hydroxy, C<sub>1-6</sub>-alkyl, C<sub>1-6</sub>-haloalkyl and C<sub>1-6</sub>-alkoxy;

wherein R<sup>1</sup> is selected from unsubstituted or substituted

aryl,  
 5-6 membered heteroaryl and  
 9-10 membered fused heteroaryl,

wherein substituted R<sup>1</sup> is substituted with one or more substituents selected from halo, C<sub>1-6</sub>-alkyl, optionally substituted C<sub>3-6</sub>-cycloalkyl, optionally substituted phenyl, C<sub>1-6</sub>-haloalkoxy, optionally substituted phenyloxy, benzyl, optionally substituted 5-6 membered heterocyclyl-C<sub>1</sub>-C<sub>2</sub>-alkylenyl, optionally substituted heteroaryl, optionally substituted heteroaryloxy, C<sub>1-6</sub>-haloalkyl, and C<sub>1-6</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,  
 halo,  
 C<sub>1-6</sub>-alkyl,  
 C<sub>1-6</sub>-haloalkyl,  
 C<sub>1-6</sub>-alkoxy,  
 C<sub>1-6</sub>-haloalkoxy,  
 C<sub>1-6</sub>-carboxyalkyl,  
 unsubstituted or substituted aryl and  
 unsubstituted or substituted 5-6 membered heteroaryl; and

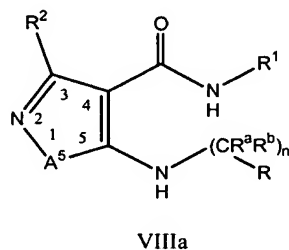
wherein R<sup>6</sup> is H or C<sub>1-6</sub>-alkyl;

and pharmaceutically acceptable isomers and salts thereof.

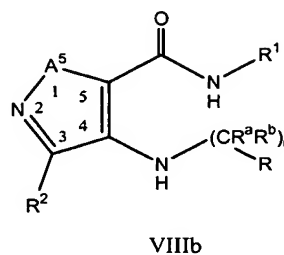
Claim 28 (withdrawn): Compound of Claim 27 wherein R<sup>a</sup> and R<sup>b</sup> are H;  
 wherein n is 1-2;  
 wherein R is selected from 4-pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, indazolyl, quinolyl, isoquinolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy and ethoxy;  
 wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, naphthyl, isoquinolyl, quinolyl, pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, naphthyridinyl, quinozaliny, tetrahydroquinoliny, indazolyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, or benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, cyclohexyl, phenylmethyl, morpholinylmethyl, methylpiperdinylmethyl, methylpiperazinylmethyl, ethyl, propyl, trifluoromethyl, phenyloxy, methoxy and ethoxy; and  
 wherein R<sup>2</sup> is one or more substituents independently selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy,

carboxymethyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl; and pharmaceutically acceptable salts thereof.

Claim 29 (withdrawn): Compound of Claim 1 of the Formulas



and



wherein A<sup>5</sup> is selected from S, O and NR<sup>6</sup>;

wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, C<sub>1-4</sub>-alkyl and -N(R<sup>6</sup>)<sub>2</sub>;

wherein n is 1-2;

wherein R is selected from

a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and

b) unsubstituted or substituted 9- or 10-membered fused nitrogen-containing heteroaryl,

where R is substituted with one or more substituents selected from halo, amino, hydroxy, C<sub>1-6</sub>-alkyl, C<sub>1-6</sub>-haloalkyl and C<sub>1-6</sub>-alkoxy;

wherein R<sup>1</sup> is selected from unsubstituted or substituted

aryl,

5-6 membered heteroaryl and

9-10 membered fused heteroaryl,

wherein substituted R<sup>1</sup> is substituted with one or more substituents selected from halo, C<sub>1-6</sub>-alkyl,

optionally substituted C<sub>3-6</sub>-cycloalkyl, optionally substituted phenyl, C<sub>1-6</sub>-haloalkoxy,

optionally substituted phenyloxy, benzyl, optionally substituted 5-6 membered heterocycl-yl-

C<sub>1</sub>.C<sub>2</sub>-alkylenyl, optionally substituted heteroaryl, optionally substituted heteroaryloxy, C<sub>1-6</sub>-haloalkyl, and C<sub>1-6</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,

halo,

C<sub>1-6</sub>-alkyl,

C<sub>1-6</sub>-haloalkyl,

C<sub>1-6</sub>-alkoxy,

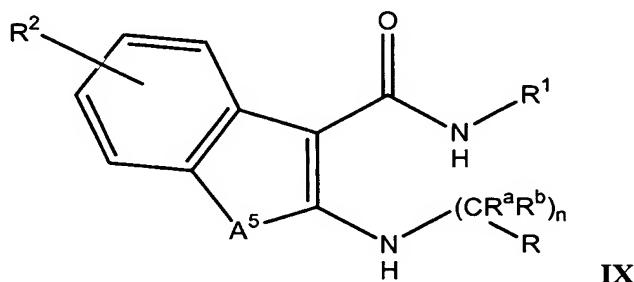
C<sub>1-6</sub>-haloalkoxy,

C<sub>1-6</sub>-carboxyalkyl,  
 unsubstituted or substituted aryl and  
 unsubstituted or substituted 5-6 membered heteroaryl; and

wherein R<sup>6</sup> is H or C<sub>1-6</sub>-alkyl;  
 and pharmaceutically acceptable isomers and salts thereof.

Claim 30 (withdrawn): Compound of Claim 29 wherein R<sup>a</sup> and R<sup>b</sup> are H;  
 wherein n is 1-2;  
 wherein R is selected from 4-pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, indazolyl,  
 quinolyl, isoquinolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or  
 substituted with one or more substituents selected  
 from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy and  
 ethoxy;  
 wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, naphthyl, isoquinolyl, quinolyl, pyridyl,  
 pyrimidinyl, pyridazinyl, indolyl, isoindolyl, naphthyridinyl, quinozaliny, tetrahydroquinolyl,  
 indazolyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, or  
 benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected  
 from chloro, fluoro, amino, hydroxy, cyclohexyl, phenylmethyl, morpholinylmethyl,  
 methylpiperdinylmethyl, methylpiperazinylmethyl, ethyl, propyl, trifluoromethyl,  
 phenyloxy, methoxy and ethoxy; and  
 wherein R<sup>2</sup> is one or more substituents independently selected from H, chloro, fluoro, bromo, amino,  
 hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy,  
 carboxymethyl, unsubstituted or substituted phenyl and unsubstituted or substituted  
 heteroaryl selected  
 from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl;  
 and pharmaceutically acceptable salts thereof.

Claim 31 (withdrawn): Compound of Claim 1 of the Formula



wherein A<sup>5</sup> is selected from S, O and NR<sup>6</sup>;  
 wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, C<sub>1-4</sub>-alkyl and -N(R<sup>6</sup>)<sub>2</sub>;

wherein n is 1-2;

wherein R is selected from

a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and

b) unsubstituted or substituted 9- or 10-membered fused nitrogen-containing heteroaryl,

where R is substituted with one or more substituents selected from halo, amino, hydroxy, C<sub>1-6</sub>-alkyl, C<sub>1-6</sub>-haloalkyl and C<sub>1-6</sub>-alkoxy;

wherein R<sup>1</sup> is selected from unsubstituted or substituted

aryl,

5-6 membered heteroaryl and

9-10 membered fused heteroaryl,

wherein substituted R<sup>1</sup> is substituted with one or more substituents selected from halo, C<sub>1-6</sub>-alkyl,

optionally substituted C<sub>3-6</sub>-cycloalkyl, optionally substituted phenyl, C<sub>1-6</sub>-haloalkoxy,

optionally substituted phenyloxy, benzyl, optionally substituted 5-6 membered heterocycl-yl-

C<sub>1-2</sub>-alkylenyl, optionally substituted heteroaryl, optionally substituted heteroaryloxy, C<sub>1-6</sub>-haloalkyl, and C<sub>1-6</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,

halo,

C<sub>1-6</sub>-alkyl,

C<sub>1-6</sub>-haloalkyl,

C<sub>1-6</sub>-alkoxy,

C<sub>1-6</sub>-haloalkoxy,

C<sub>1-6</sub>-carboxyalkyl,

unsubstituted or substituted aryl and

unsubstituted or substituted 5-6 membered heteroaryl; and

wherein R<sup>6</sup> is H or C<sub>1-6</sub>-alkyl;

and pharmaceutically acceptable isomers and salts thereof.

Claim 32 (withdrawn): Compound of Claim 31 wherein R<sup>a</sup> and R<sup>b</sup> are H;

wherein n is 1-2;

wherein R is selected from 4-pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, indazolyl,

quinolyl, isoquinolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or substituted with one or more substituents selected

from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy and ethoxy;

wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, naphthyl, isoquinolyl, quinolyl, pyridyl,

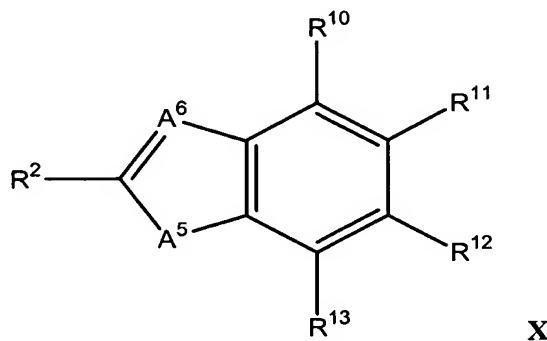
pyrimidinyl, pyridazinyl, indolyl, isoindolyl, naphthyridinyl, quinozaliny,



tetrahydroquinolinyl, indazolyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, or benzthiazolyl, where  $R^1$  is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, cyclohexyl, phenylmethyl, morpholinylmethyl, methylpiperdinylmethyl, methylpiperazinylmethyl, ethyl, propyl, trifluoromethyl, phenyloxy, methoxy and ethoxy; and

wherein  $R^2$  is one or more substituents independently selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl; and pharmaceutically acceptable salts thereof.

Claim 33 (withdrawn): Compound of Claim 1 of the Formula



wherein  $A^5$  is selected from S, O and  $NR^6$ ;

wherein  $A^6$  is selected from  $CR^2$  and N;

wherein R is selected from

a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and

b) unsubstituted or substituted 9- or 10-membered fused nitrogen-containing heteroaryl,

where R is substituted with one or more substituents selected from halo, amino, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl and  $C_{1-6}$ -alkoxy;

wherein  $R^1$  is selected from unsubstituted or substituted

aryl,

5-6 membered heteroaryl and

9-10 membered fused heteroaryl,

wherein substituted  $R^1$  is substituted with one or more substituents selected from halo,  $C_{1-6}$ -alkyl,

optionally substituted  $C_{3-6}$ -cycloalkyl, optionally substituted phenyl,  $C_{1-6}$ -haloalkoxy,

optionally substituted phenyloxy, benzyl, optionally substituted 5-6 membered heterocyclyl-

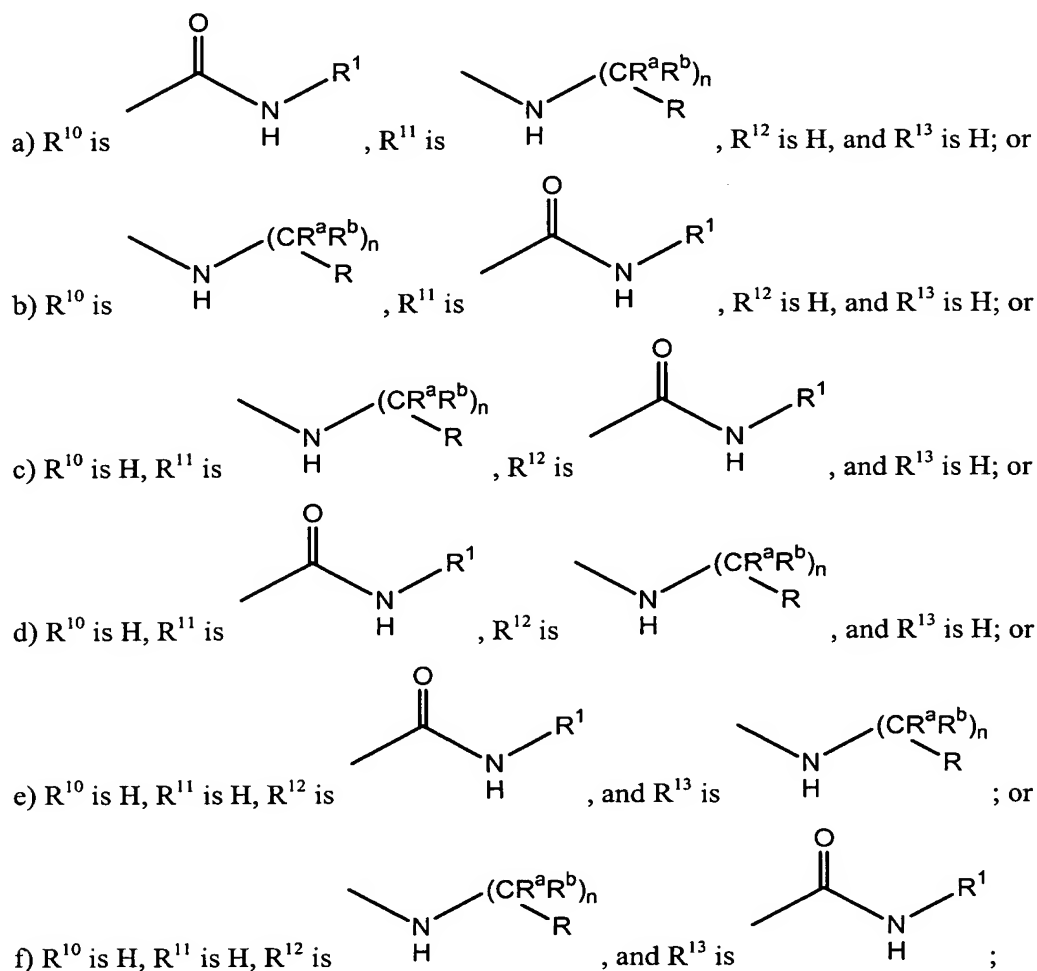
C<sub>1</sub>-C<sub>2</sub>-alkylenyl, optionally substituted heteroaryl, optionally substituted heteroaryloxy, C<sub>1-6</sub>-haloalkyl, and C<sub>1-6</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,  
halo,  
C<sub>1-6</sub>-alkyl,  
C<sub>1-6</sub>-haloalkyl,  
C<sub>1-6</sub>-alkoxy,  
C<sub>1-6</sub>-haloalkoxy,  
C<sub>1-6</sub>-carboxyalkyl,  
unsubstituted or substituted aryl and  
unsubstituted or substituted 5-6 membered heteroaryl; and

wherein R<sup>6</sup> is H or C<sub>1-6</sub>-alkyl;

wherein



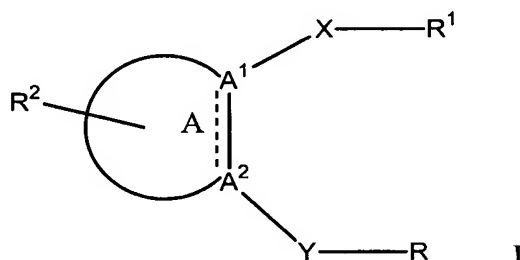
wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, C<sub>1-4</sub>-alkyl and -N(R<sup>6</sup>)<sub>2</sub>; and  
wherein n is 1-2;

and pharmaceutically acceptable isomers and salts thereof.

Claim 34 (withdrawn): Compound of Claim 33 wherein  $R^a$  and  $R^b$  are H;  
 wherein n is 1-2;  
 wherein R is selected from 4-pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, indazolyl, quinolyl, isoquinolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy and ethoxy;  
 wherein  $R^1$  is selected from phenyl, tetrahydronaphthyl, naphthyl, isoquinolyl, quinolyl, pyridyl, pyrimidinyl, pyridazinyl, indolyl, isoindolyl, naphthyridinyl, quinozaliny, tetrahydroquinoliny, indazolyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl, or benzthiazolyl, where  $R^1$  is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, cyclohexyl, phenylmethyl, morpholinylmethyl, methylpiperdinylmethyl, methylpiperazinylmethyl, ethyl, propyl, trifluoromethyl, phenyloxy, methoxy and ethoxy; and  
 wherein  $R^2$  is one or more substituents independently selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl;  
 and pharmaceutically acceptable salts thereof.

Claim 35 (canceled).

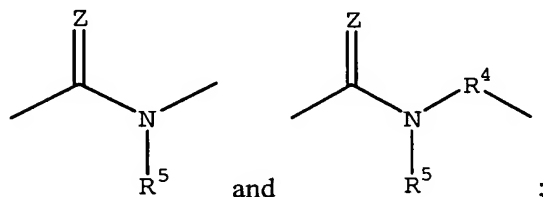
Claim 36 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Formula I



wherein each of  $A^1$  and  $A^2$  is independently C or N;  
 wherein ring A is selected from  
 a) 5- or 6-membered partially saturated heterocycl,

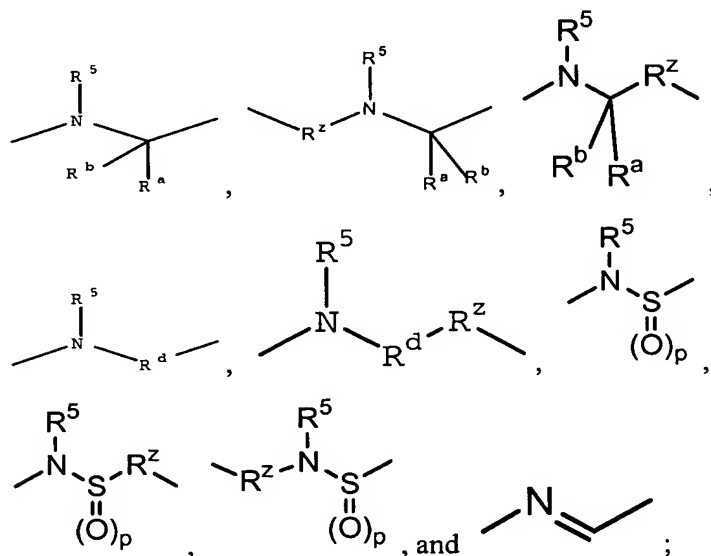
- b) 5- or 6-membered heteroaryl,
- c) 9- or 10-membered fused partially saturated heterocyclyl,
- d) 9-, 10- or 11-membered fused heteroaryl;
- e) naphthyl, and
- f) 4-, 5- or 6- membered cycloalkenyl;

wherein X is selected from



wherein Z is oxygen or sulfur;

wherein Y is selected from



wherein p is 0 to 2,

wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, cyano, -NHR<sup>6</sup> and C<sub>1-4</sub>-alkyl substituted with R<sup>2</sup>, or wherein R<sup>a</sup> and R<sup>b</sup> together form C<sub>3</sub>-C<sub>6</sub> cycloalkyl;

wherein R<sup>z</sup> is selected from C<sub>1</sub>-C<sub>4</sub> alkylenyl, where one of the CH<sub>2</sub> groups may be substituted with an oxygen atom or an -NH-;

wherein R<sup>d</sup> is cycloalkyl;

wherein R is selected from

- a) substituted or unsubstituted 5-6 membered heterocyclyl, and
- b) substituted or unsubstituted fused 9-, 10- or 11-membered heterocyclyl;

wherein substituted R is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, cycloalkyl, optionally substituted 5-6 membered

heterocyclyl, optionally substituted phenyl, lower alkyl substituted with  $R^2$ , cyano, nitro, lower alkenyl and lower alkynyl;

wherein  $R^1$  is selected from

- a) substituted or unsubstituted 6-10 membered aryl,
- b) substituted or unsubstituted 5-6 membered heterocyclyl,
- c) substituted or unsubstituted 9-11 membered fused heterocyclyl,
- d) cycloalkyl, and
- e) cycloalkenyl,

wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-NH(C_1-C_4 \text{ alkylenyl}R^{14})$ ,  $-SO_2R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, lower alkyl substituted with  $R^2$ , cyano, nitro, lower alkenyl and lower alkynyl;

wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,  $-COR^3$ ,  $-CONR^3R^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted phenylalkylenyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted heteroarylalkylenyl, optionally substituted phenyl, lower alkyl, cyano, lower hydroxyalkyl, lower carboxyalkyl, nitro, lower alkenyl, lower alkynyl, lower aminoalkyl, lower alkylaminoalkyl and lower haloalkyl;

wherein  $R^3$  is selected from H, lower alkyl, phenyl, 5-6 membered heterocyclyl,  $C_3-C_6$  cycloalkyl, and lower haloalkyl;

wherein  $R^4$  is independently selected from  $C_2-C_4$  alkylenyl,  $C_2-C_4$  alkenylenyl and  $C_2-C_4$  alkynylenyl, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ;

wherein  $R^5$  is selected from H, lower alkyl, phenyl and lower aralkyl; and

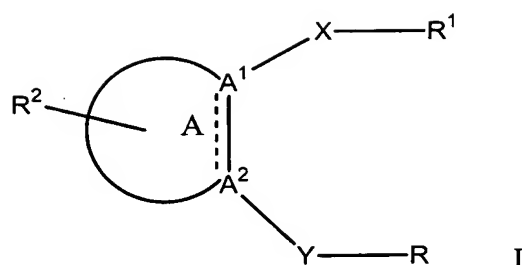
wherein  $R^6$  is selected from H or  $C_{1-6}$ -alkyl;

wherein  $R^{14}$  is selected from H, phenyl, 5-6 membered heterocyclyl and  $C_3-C_6$  cycloalkyl; and pharmaceutically acceptable salts thereof;

provided A is not naphthyl when X is  $-C(O)NH-$  and when  $R^1$  is phenyl when Y is  $-NCH_2-$  and when R is 4-pyridyl; and further provided R is not unsubstituted 2-thienyl, 2-pyridyl or 3-pyridyl when Y is  $-NHCH_2-$ .

Claim 37 (withdrawn): The method of Claim 36 comprising a combination with a compound selected from antibiotic-type agents, alkylating agents, antimetabolite agents, hormonal agents, immunological agents, interferon-type agents and miscellaneous agents.

Claim 38 (withdrawn): A method of treating angiogenesis in a subject, said method comprising administering an effective amount of a compound as in any of Formula I

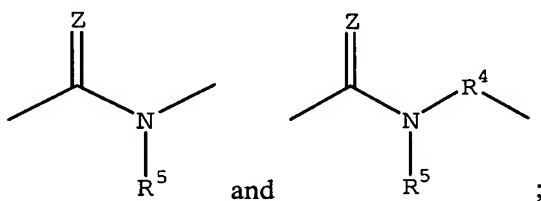


wherein each of  $A^1$  and  $A^2$  is independently C or N;

wherein ring A is selected from

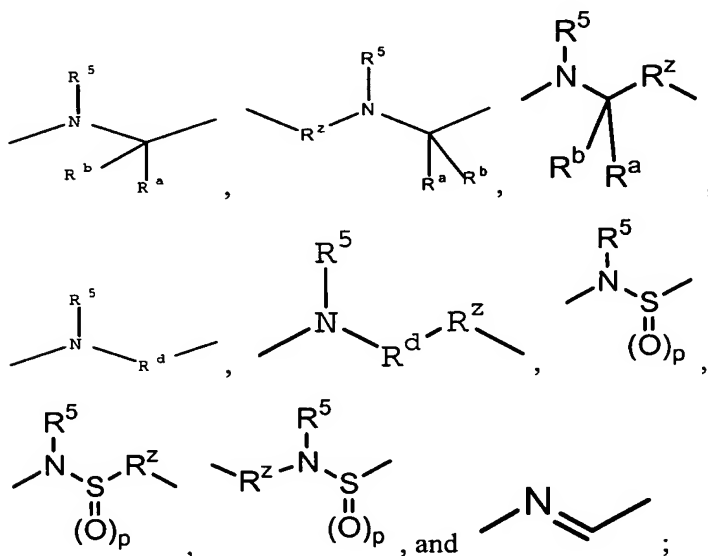
- a) 5- or 6-membered partially saturated heterocyclyl,
- b) 5- or 6-membered heteroaryl,
- c) 9- or 10-membered fused partially saturated heterocyclyl,
- d) 9-, 10- or 11-membered fused heteroaryl;
- e) naphthyl, and
- f) 4-, 5- or 6- membered cycloalkenyl;

wherein X is selected from



wherein Z is oxygen or sulfur;

wherein Y is selected from



wherein p is 0 to 2,

wherein  $R^a$  and  $R^b$  are independently selected from H, halo, cyano,  $-NHR^6$  and  $C_{1-4}$ -alkyl substituted with  $R^2$ , or wherein  $R^a$  and  $R^b$  together form  $C_3$ - $C_6$  cycloalkyl;

wherein  $R^z$  is selected from  $C_1$ - $C_4$  alkylenyl, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ;

wherein  $R^d$  is cycloalkyl;

wherein R is selected from

a) substituted or unsubstituted 5-6 membered heterocyclyl, and

b) substituted or unsubstituted fused 9-, 10- or 11-membered heterocyclyl;

wherein substituted R is substituted with one or more substituents independently selected

from halo,  $-OR^3$ ,  $-SR^3$ ,  $-SO_2R^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ , -

$NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered

heterocyclyl, optionally substituted phenyl, lower alkyl substituted with  $R^2$ , cyano, nitro,

lower alkenyl and lower alkynyl;

wherein  $R^1$  is selected from

a) substituted or unsubstituted 6-10 membered aryl,

b) substituted or unsubstituted 5-6 membered heterocyclyl,

c) substituted or unsubstituted 9-11 membered fused heterocyclyl,

d) cycloalkyl, and

e) cycloalkenyl,

wherein substituted  $R^1$  is substituted with one or more substituents independently selected

from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-NH(C_1-C_4 \text{ alkylenyl}R^{14})$ , -

$SO_2R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , optionally substituted cycloalkyl,

optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, lower

alkyl substituted with  $R^2$ , cyano, nitro, lower alkenyl and lower alkynyl;

wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,

$-COR^3$ ,  $-CONR^3R^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally

substituted phenylalkylenyl, optionally substituted 5-6 membered heterocyclyl, optionally

substituted heteroarylalkylenyl, optionally substituted phenyl, lower alkyl, cyano, lower

hydroxyalkyl, lower carboxyalkyl, nitro, lower alkenyl, lower alkynyl, lower aminoalkyl, lower

alkylaminoalkyl and lower haloalkyl;

wherein  $R^3$  is selected from H, lower alkyl, phenyl, 5-6 membered heterocyclyl,  $C_3$ - $C_6$  cycloalkyl, and lower haloalkyl;

wherein  $R^4$  is independently selected from  $C_2$ - $C_4$  alkylenyl,  $C_2$ - $C_4$  alkenylenyl and  $C_2$ - $C_4$  alkynylenyl,

where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ;

wherein  $R^5$  is selected from H, lower alkyl, phenyl and lower aralkyl; and

wherein  $R^6$  is selected from H or  $C_{1-6}$ -alkyl;

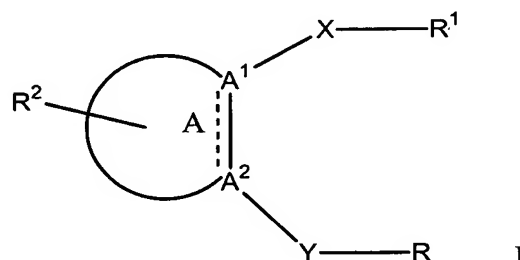
wherein  $R^{14}$  is selected from H, phenyl, 5-6 membered heterocyclyl and  $C_3$ - $C_6$  cycloalkyl;

and pharmaceutically acceptable salts thereof;

provided A is not naphthyl when X is -C(O)NH- and when R<sup>1</sup> is phenyl when Y is -NCH<sub>2</sub>- and when R is 4-pyridyl; and further provided R is not unsubstituted 2-thienyl, 2-pyridyl or 3-pyridyl when Y is -NHCH<sub>2</sub>-.

Claim 39 (canceled).

Claim 40 (withdrawn): A method of treating KDR-related disorders in a mammal, said method comprising administering an effective amount of a compound of Formula I

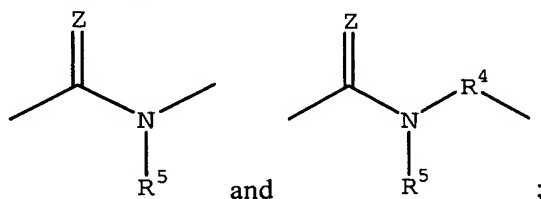


wherein each of A<sup>1</sup> and A<sup>2</sup> is independently C or N;

wherein ring A is selected from

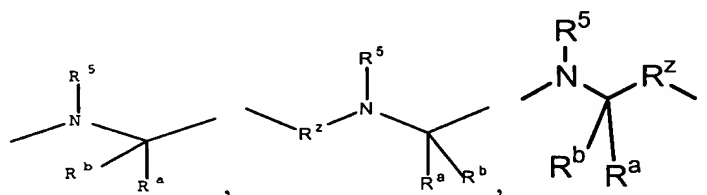
- a) 5- or 6-membered partially saturated heterocyclyl,
- b) 5- or 6-membered heteroaryl,
- c) 9- or 10-membered fused partially saturated heterocyclyl,
- d) 9-, 10- or 11-membered fused heteroaryl;
- e) naphthyl, and
- f) 4-, 5- or 6- membered cycloalkenyl;

wherein X is selected from

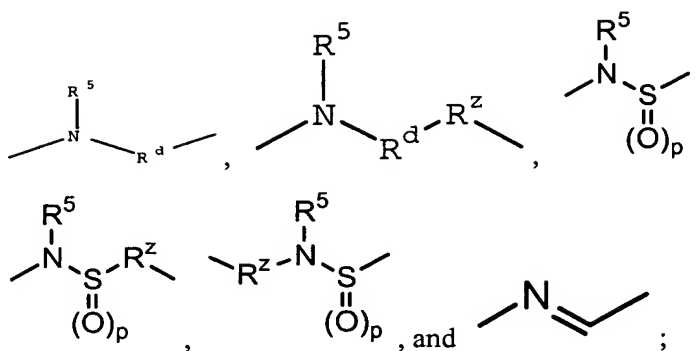


wherein Z is oxygen or sulfur;

wherein Y is selected from







wherein  $p$  is 0 to 2,

wherein  $R^a$  and  $R^b$  are independently selected from H, halo, cyano,  $-NHR^6$  and  $C_{1-4}$ -alkyl substituted with  $R^2$ , or wherein  $R^a$  and  $R^b$  together form  $C_3$ - $C_6$  cycloalkyl;

wherein  $R^z$  is selected from  $C_1$ - $C_4$  alkylene, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH$ -;

wherein  $R^d$  is cycloalkyl;

wherein  $R$  is selected from

- substituted or unsubstituted 5-6 membered heterocyclyl, and
- substituted or unsubstituted fused 9-, 10- or 11-membered heterocyclyl;

wherein substituted  $R$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-SO_2R^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, lower alkyl substituted with  $R^2$ , cyano, nitro, lower alkenyl and lower alkynyl;

wherein  $R^1$  is selected from

- substituted or unsubstituted 6-10 membered aryl,
- substituted or unsubstituted 5-6 membered heterocyclyl,
- substituted or unsubstituted 9-11 membered fused heterocyclyl,
- cycloalkyl, and
- cycloalkenyl,

wherein substituted  $R^1$  is substituted with one or more substituents independently selected from halo,  $-OR^3$ ,  $-SR^3$ ,  $-CO_2R^3$ ,  $-CONR^3R^3$ ,  $-COR^3$ ,  $-NR^3R^3$ ,  $-NH(C_1$ - $C_4$  alkylene $R^{14})$ ,  $-SO_2R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , optionally substituted cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, lower alkyl substituted with  $R^2$ , cyano, nitro, lower alkenyl and lower alkynyl;

wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,  $-COR^3$ ,  $-CONR^3R^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted phenylalkylene, optionally substituted 5-6 membered heterocyclyl, optionally substituted heteroarylalkylene, optionally substituted phenyl, lower alkyl, cyano, lower

hydroxyalkyl, lower carboxyalkyl, nitro, lower alkenyl, lower alkynyl, lower aminoalkyl, lower alkylaminoalkyl and lower haloalkyl;

wherein  $R^3$  is selected from H, lower alkyl, phenyl, 5-6 membered heterocyclyl,  $C_3$ - $C_6$  cycloalkyl, and lower haloalkyl;

wherein  $R^4$  is independently selected from  $C_2$ - $C_4$  alkylenyl,  $C_2$ - $C_4$  alkenylenyl and  $C_2$ - $C_4$  alkynylenyl,

where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ;

wherein  $R^5$  is selected from H, lower alkyl, phenyl and lower aralkyl; and

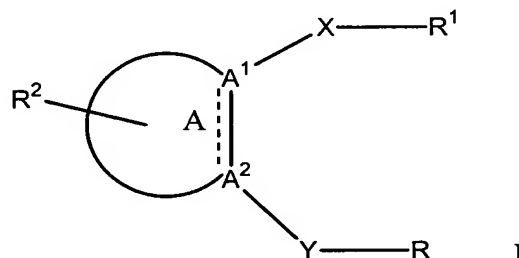
wherein  $R^6$  is selected from H or  $C_{1-6}$ -alkyl;

wherein  $R^{14}$  is selected from H, phenyl, 5-6 membered heterocyclyl and  $C_3$ - $C_6$  cycloalkyl;

and pharmaceutically acceptable salts thereof;

provided A is not naphthyl when X is  $-C(O)NH-$  and when  $R^1$  is phenyl when Y is  $-NCH_2-$  and when R is 4-pyridyl; and further provided R is not unsubstituted 2-thienyl, 2-pyridyl or 3-pyridyl when Y is  $-NHCH_2-$ .

Claim 41 (withdrawn): A method of treating proliferative disorders in a mammal, said method comprising administering an effective amount of a compound of Formula I

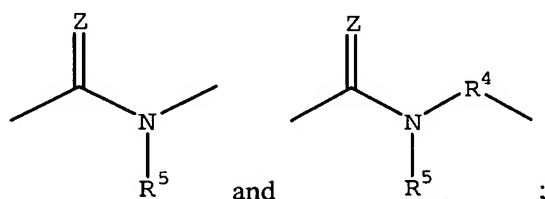


wherein each of  $A^1$  and  $A^2$  is independently C or N;

wherein ring A is selected from

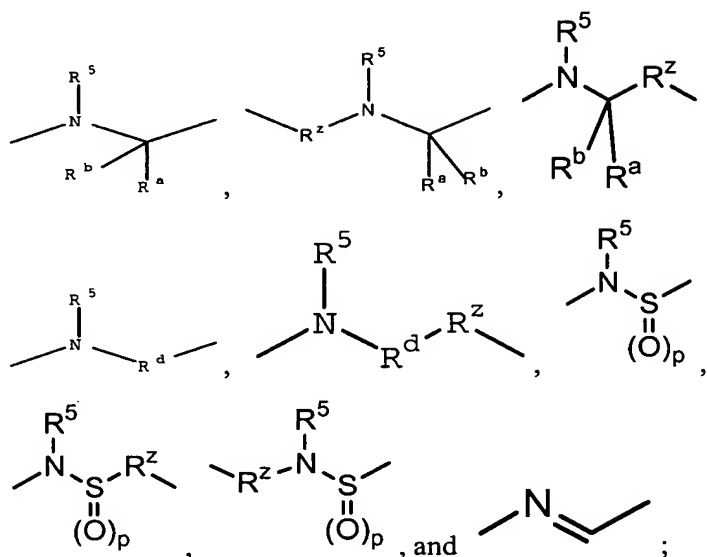
- a) 5- or 6-membered partially saturated heterocyclyl,
- b) 5- or 6-membered heteroaryl,
- c) 9- or 10-membered fused partially saturated heterocyclyl,
- d) 9-, 10- or 11-membered fused heteroaryl;
- e) naphthyl, and
- f) 4-, 5- or 6- membered cycloalkenyl;

wherein X is selected from



wherein Z is oxygen or sulfur;

wherein Y is selected from



wherein p is 0 to 2,

wherein R<sup>a</sup> and R<sup>b</sup> are independently selected from H, halo, cyano, -NHR<sup>6</sup> and C<sub>1-4</sub>-alkyl substituted with R<sup>2</sup>, or wherein R<sup>a</sup> and R<sup>b</sup> together form C<sub>3-6</sub> cycloalkyl;

wherein R<sup>z</sup> is selected from C<sub>1-4</sub> alkylenyl, where one of the CH<sub>2</sub> groups may be substituted with an oxygen atom or an -NH-;

wherein R<sup>d</sup> is cycloalkyl;

wherein R is selected from

- substituted or unsubstituted 5-6 membered heterocyclyl, and
- substituted or unsubstituted fused 9-, 10- or 11-membered heterocyclyl;

wherein substituted R is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -SO<sub>2</sub>R<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, cycloalkyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, lower alkyl substituted with R<sup>2</sup>, cyano, nitro, lower alkenyl and lower alkynyl;

wherein R<sup>1</sup> is selected from

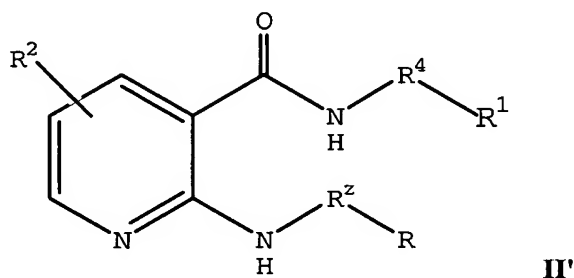
- substituted or unsubstituted 6-10 membered aryl,
- substituted or unsubstituted 5-6 membered heterocyclyl,
- substituted or unsubstituted 9-11 membered fused heterocyclyl,
- cycloalkyl, and
- cycloalkenyl,

wherein substituted R<sup>1</sup> is substituted with one or more substituents independently selected from halo, -OR<sup>3</sup>, -SR<sup>3</sup>, -CO<sub>2</sub>R<sup>3</sup>, -CONR<sup>3</sup>R<sup>3</sup>, -COR<sup>3</sup>, -NR<sup>3</sup>R<sup>3</sup>, -NH(C<sub>1-4</sub> alkylenylR<sup>14</sup>), -SO<sub>2</sub>R<sup>3</sup>, -SO<sub>2</sub>NR<sup>3</sup>R<sup>3</sup>, -NR<sup>3</sup>C(O)OR<sup>3</sup>, -NR<sup>3</sup>C(O)R<sup>3</sup>, optionally substituted cycloalkyl,

optionally substituted 5-6 membered heterocyclyl, optionally substituted phenyl, lower alkyl substituted with  $R^2$ , cyano, nitro, lower alkenyl and lower alkynyl;  
 wherein  $R^2$  is one or more substituents independently selected from H, halo,  $-OR^3$ , oxo,  $-SR^3$ ,  $-CO_2R^3$ ,  $-COR^3$ ,  $-CONR^3R^3$ ,  $-NR^3R^3$ ,  $-SO_2NR^3R^3$ ,  $-NR^3C(O)OR^3$ ,  $-NR^3C(O)R^3$ , cycloalkyl, optionally substituted phenylalkylenyl, optionally substituted 5-6 membered heterocyclyl, optionally substituted heteroarylalkylenyl, optionally substituted phenyl, lower alkyl, cyano, lower hydroxyalkyl, lower carboxyalkyl, nitro, lower alkenyl, lower alkynyl, lower aminoalkyl, lower alkylaminoalkyl and lower haloalkyl;  
 wherein  $R^3$  is selected from H, lower alkyl, phenyl, 5-6 membered heterocyclyl,  $C_3$ - $C_6$  cycloalkyl, and lower haloalkyl;  
 wherein  $R^4$  is independently selected from  $C_2$ - $C_4$  alkylenyl,  $C_2$ - $C_4$  alkenylenyl and  $C_2$ - $C_4$  alkynylenyl, where one of the  $CH_2$  groups may be substituted with an oxygen atom or an  $-NH-$ ;  
 wherein  $R^5$  is selected from H, lower alkyl, phenyl and lower aralkyl; and  
 wherein  $R^6$  is selected from H or  $C_{1-6}$ -alkyl;  
 wherein  $R^{14}$  is selected from H, phenyl, 5-6 membered heterocyclyl and  $C_3$ - $C_6$  cycloalkyl;  
 and pharmaceutically acceptable salts thereof;  
 provided A is not naphthyl when X is  $-C(O)NH-$  and when  $R^1$  is phenyl when Y is  $-NCH_2-$  and when R is 4-pyridyl; and further provided R is not unsubstituted 2-thienyl, 2-pyridyl or 3-pyridyl when Y is  $-NHCH_2-$ .

Claim 42 (withdrawn): Method of Claim 41 wherein the disorder is inflammation or an inflammation-related disorder.

Claim 43 (currently amended): A compound of ~~Claim 1~~ having Formula II'



wherein R is selected from

- a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and
  - b) unsubstituted or substituted 9- or 10-membered fused heteroaryl,
- where substituted R is substituted with one or more substituents selected from halo, amino, oxo, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl,  $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkylamino,

optionally substituted heterocyclyl-C<sub>1-6</sub>-alkyl, C<sub>1-6</sub>-alkylamino-C<sub>2-4</sub>-alkynyl, C<sub>1-6</sub>-alkylamino-C<sub>1-6</sub>-alkoxy, C<sub>1-6</sub>-alkylamino-C<sub>1-6</sub>-alkoxy-C<sub>1-6</sub>-alkoxy, and optionally substituted heterocyclyl-C<sub>2-4</sub>-alkynyl;

wherein R<sup>1</sup> is selected from unsubstituted or substituted

aryl,

cycloalkyl,

5-6 membered heteroaryl and

9-10 membered bicyclic and 13-14 membered tricyclic heterocyclyl,

wherein substituted R<sup>1</sup> is substituted with one or more substituents selected from halo, C<sub>1-6</sub>-alkyl,

optionally substituted C<sub>3-6</sub>-cycloalkyl, optionally substituted phenyl, optionally substituted

phenyl-C<sub>1</sub>-C<sub>4</sub>-alkylenyl, C<sub>1-2</sub>-haloalkoxy, optionally substituted 4-6 membered heterocyclyl-

C<sub>1</sub>-C<sub>4</sub>-alkyl, optionally substituted 4-6 membered heterocyclyl-C<sub>2</sub>-C<sub>4</sub>-alkenyl, optionally

substituted 4-6 membered heterocyclyl, optionally substituted phenyloxy, optionally

substituted 4-6 membered heterocycliloxy, optionally substituted 4-6 membered

heterocyclyl-C<sub>1</sub>-C<sub>4</sub>-alkoxy, optionally substituted 4-6 membered heterocyclylsulfonyl,

optionally substituted 4-6 membered heterocyclylamino, optionally substituted 4-6 membered

heterocyclylcarbonyl, optionally substituted 5-6 membered heterocyclyl-C<sub>1-4</sub>-alkylcarbonyl,

C<sub>1-2</sub>-haloalkyl, C<sub>1-4</sub>-aminoalkyl, nitro, amino, hydroxy, oxo, cyano, -NHC(O)NH<sub>2</sub>,

alkylcarbonylamino, aminosulfonyl, C<sub>1-2</sub>-alkylsulfonyl, halosulfonyl, C<sub>1-4</sub>-alkylcarbonyl, C<sub>1-3</sub>-

alkylamino-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy,

C<sub>1-4</sub>-alkoxycarbonyl, C<sub>1-4</sub>-alkoxycarbonylamino-C<sub>1-4</sub>-alkyl, C<sub>1-4</sub>-hydroxyalkyl,

and C<sub>1-4</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,

halo,

hydroxy,

amino,

C<sub>1-6</sub>-alkyl,

C<sub>1-6</sub>-haloalkyl,

C<sub>1-6</sub>-alkoxy,

C<sub>1-2</sub>-alkylamino,

aminosulfonyl,

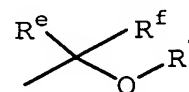
C<sub>3-6</sub>-cycloalkyl,

cyano,

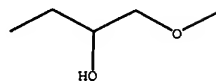
C<sub>1-2</sub>-hydroxyalkyl,

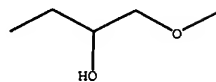
nitro,

C<sub>2-3</sub>-alkenyl,



$C_{2-3}$ -alkynyl,  
 $C_{1-6}$ -haloalkoxy,  
 $C_{1-6}$ -carboxyalkyl,  
 5-6-membered heterocyclyl- $C_{1-6}$ -alkylamino,  
~~unsubstituted or substituted phenyl and~~  
 unsubstituted or substituted 5-6 membered heterocyclyl;



wherein  $R^4$  is selected from a direct bond,  $C_{1-4}$ -alkyl, and  ;  
 wherein  $R^z$  is selected from  $C_{1-2}$ -alkyl,  $C_{2-6}$ -branched alkyl,  $C_{2-4}$ -branched haloalkyl, amino- $C_{1-4}$ -alkyl and  $C_{1-2}$ -alkylamino- $C_{1-2}$ -alkyl;  
 wherein  $R^e$  and  $R^f$  are independently selected from H and  $C_{1-2}$ -haloalkyl; and  
 wherein  $R^7$  is selected from H,  $C_{1-3}$ -alkyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-3}$ -alkyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted 4-6 membered heterocyclyl- $C_{1-3}$ -alkyl,  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkyl,  $C_{1-3}$ -alkoxy- $C_{1-2}$ -alkyl and  $C_{1-3}$ -alkoxy- $C_{1-3}$ -alkoxy- $C_{1-3}$ -alkyl;  
 1) provided  $R^2$  is not H, or  
 2) provided  $R^1$  is not heteroaryl or aryl, or  
 3) provided R is substituted with optionally substituted heterocyclyl- $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkylamino, optionally substituted heterocyclyl- $C_{1-6}$ -alkyl,  $C_{1-6}$ -alkylamino- $C_{2-4}$ -alkynyl,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy- $C_{1-6}$ -alkoxy, or optionally substituted heterocyclyl- $C_{2-4}$ -alkynyl, or  
 4) provided  $R^1$  is substituted with optionally substituted phenyloxy, optionally substituted 5-6 membered heterocyclyloxy, optionally substituted 5-6 membered heterocyclylsulfonyl, optionally substituted 5-6 membered heterocyclylamino, optionally substituted 5-6 membered heterocyclylcarbonyl, optionally substituted 5-6 membered heterocyclyl- $C_{1-4}$ -alkylcarbonyl,  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkoxy, or  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkoxy- $C_{1-3}$ -alkoxy;  
 further provided R is not 3-pyridyl when  $R^s$   $R^z$  is  $CH_2$ ;  
further provided  $R^1$  is not 4-[3-(3-pyridyl)-5-(trifluoromethyl)-1H-pyrazol-1-yl]phenyl when  $R^z$  is  $CH_2$  and when R is 4-pyridyl;

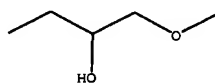
and pharmaceutically acceptable isomers and derivatives thereof.

Claim 44 (currently amended): Compound of Claim 43 wherein R is selected from 4-pyridyl, 3-pyridyl, 2-pyridyl, pyrimidinyl, triazolyl, pyridazinyl, indolyl, isoindolyl, indazolyl, quinolyl, isoquinolyl, benzotriazolyl, 2,3-dihydrobenzofuryl, 2-oxo-1,2-dihydroquinol-7-yl, naphthyridinyl and quinoxalinyl, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, dimethylaminopropynyl, 1-methylpiperidinylmethoxy, dimethylaminoethoxyethoxy, methoxy and ethoxy; wherein  $R^1$  is selected

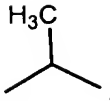
from phenyl, tetrahydronaphthyl, indanyl, indenyl, naphthyl, cyclohexyl, isoxazolyl, pyrazolyl, thiazolyl, thiadiazolyl, thienyl, pyridyl, pyrimidinyl, pyridazinyl, 1,2-dihydroquinolyl, 1,2,3,4-tetrahydro-isoquinolyl, isoquinolyl, quinolyl, indolyl, isoindolyl, 2,3-dihydro-1H-indolyl, naphthyridinyl, quinoxalinyl, benzo[d]isothiazolyl, 2,3,4,4a,9,9a-hexahydro-1H-3-aza-fluorenyl, 5,6,7-trihydro-1,2,4-triazolo[3,4-a]isoquinolyl, tetrahydroquinolinyl, indazolyl, 2,1,3-benzothiadiazolyl, benzodioxanyl, benzothienyl, benzofuryl, dihydro-benzimidazolyl, benzimidazolyl, benzoxazolyl and benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from bromo, chloro, fluoro, iodo, nitro, amino, cyano, aminoethyl, Boc-aminoethyl, hydroxy, oxo, aminosulfonyl, 4-methylpiperazinylsulfonyl, cyclohexyl, phenyl, phenylmethyl, morpholinylmethyl, 1-methylpiperazin-4-ylmethyl, 1-methylpiperazin-4-ylpropyl, morpholinylpropyl, piperidin-1-ylmethyl, 1-methylpiperidin-4-ylmethyl, 2-methyl-2-(1-methylpiperidin-4-yl)ethyl, morpholinylethyl, 1-(4-morpholinyl)-2,2-dimethylpropyl, piperidin-4-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-1-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-4-ylmethyl, 1-Boc-piperidin-4-ylmethyl, piperidin-4-ylpropyl, 1-Boc-piperidin-4-ylpropyl, piperidin-1-ylpropyl, pyrrolidin-1-ylpropyl, pyrrolidin-2-ylpropyl, 1-Boc-pyrrolidin-2-ylpropyl, pyrrolidin-1-ylmethyl, pyrrolidin-2-ylmethyl, 1-Boc-pyrrolidin-2-ylmethyl, pyrrolidinylpropenyl, pyrrolidinylbutenyl, fluorosulfonyl, methylsulfonyl, methylcarbonyl, Boc, piperidin-1-ylmethylcarbonyl, 4-methylpiperazin-1-ylcarbonylethyl, methoxycarbonyl, aminomethylcarbonyl, dimethylaminomethylcarbonyl, 3-ethoxycarbonyl-2-methyl-fur-5-yl, 4-methylpiperazin-1-yl, 4-methyl-1-piperidyl, 1-Boc-4-piperidyl, piperidin-4-yl, 1-methylpiperidin-4-yl, 1-methyl-(1,2,3,6-tetrahydropyridyl), imidazolyl, morpholinyl, 4-trifluoromethyl-1-piperidinyl, hydroxybutyl, methyl, ethyl, propyl, isopropyl, butyl, tert-butyl, sec-butyl, trifluoromethyl, pentafluoroethyl, nonafluorobutyl, dimethylaminopropyl, 1,1-di(trifluoromethyl)-1-hydroxymethyl, 1,1-di(trifluoromethyl)-1-(piperidinylethoxy)methyl, 1,1-di(trifluoromethyl)-1-(methoxyethoxyethoxy)methyl, 1-hydroxyethyl, 2-hydroxyethyl, trifluoromethoxy, 1-aminoethyl, 2-aminoethyl, 1-(N-isopropylamino)ethyl, 2-(N-isopropylamino)ethyl, dimethylaminoethoxy, 4-chlorophenoxy, phenyloxy, azetidin-3-ylmethoxy, 1-Boc-azetidin-3-ylmethoxy, pyrrol-2-ylmethoxy, 1-Boc-pyrrol-2-ylmethoxy, pyrrol-1-ylmethoxy, 1-methyl-pyrrol-2-ylmethoxy, 1-isopropyl-pyrrol-2-ylmethoxy, 1-Boc-piperdin-4-ylmethoxy, piperdin-4-ylmethoxy, 1-methylpiperdin-4-yloxy, isopropoxy, methoxy and ethoxy; wherein R<sup>2</sup> is selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, oxo, dimethylamino, aminosulfonyl, cyclopropyl, cyano, hydroxymethyl, nitro, propenyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, morpholinylethylamino, propynyl, ~~unsubstituted or substituted phenyl~~ and unsubstituted or substituted heteroaryl selected from thienyl,

furanyl, pyridyl, imidazolyl, and pyrazolyl;

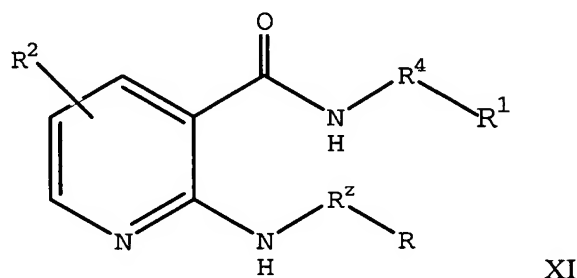
wherein R<sup>4</sup> is selected from a direct bond, ethyl, butyl, and



; and

wherein  $R^z$  is selected from methylenyl, ethylenyl, , and aminoethylenyl; and pharmaceutically acceptable derivatives thereof.

Claim 45 (currently amended): A compound of Claim ~~43~~ having Formula XI



wherein R is selected from

- a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and
- b) unsubstituted or substituted 9- or 10-membered fused heteroaryl,

where substituted R is substituted with one or more substituents selected from halo, amino, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl,  $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkylamino, optionally substituted heterocyclyl- $C_{1-6}$ -alkyl,  $C_{1-6}$ -alkylamino- $C_{2-4}$ -alkynyl,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy- $C_{1-6}$ -alkoxy, and optionally substituted heterocyclyl- $C_{2-4}$ -alkynyl;

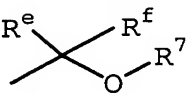
wherein  $R^1$  is a ring selected from unsubstituted or substituted

- 4-6 membered saturated or partially un-saturated monocyclic heterocyclyl,
- 9-10 membered saturated or partially un-saturated bicyclic heterocyclyl, and
- 13-14 membered saturated or partially un-saturated tricyclic heterocyclyl,

wherein substituted  $R^1$  is substituted with one or more substituents selected from halo,  $C_{1-6}$ -alkyl, optionally substituted  $C_{3-6}$ -cycloalkyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-4}$ -alkylenyl,  $C_{1-2}$ -haloalkoxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkyl, optionally substituted 4-6 membered heterocyclyl- $C_{2-4}$ -alkenyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted phenyloxy, optionally substituted 4-6 membered heterocyclyloxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkoxy, optionally substituted 4-6 membered heterocyclylsulfonyl, optionally substituted 4-6 membered heterocyclylamino, optionally substituted 4-6 membered heterocyclylcarbonyl, optionally substituted 5-6 membered heterocyclyl- $C_{1-4}$ -alkylcarbonyl,  $C_{1-2}$ -haloalkyl,  $C_{1-4}$ -aminoalkyl, nitro, amino, hydroxy, oxo, cyano, aminosulfonyl,  $C_{1-2}$ -alkylsulfonyl, halosulfonyl,  $C_{1-4}$ -alkylcarbonyl,  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkyl,  $C_{1-3}$ -alkylamino-



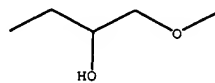
C<sub>1-3</sub>-alkoxy, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy, C<sub>1-4</sub>-alkoxycarbonyl, C<sub>1-4</sub>-

alkoxycarbonylamino-C<sub>1-4</sub>-alkyl, C<sub>1-4</sub>-hydroxyalkyl,  and C<sub>1-4</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,  
halo,  
hydroxy,  
amino,  
C<sub>1-6</sub>-alkyl,  
C<sub>1-6</sub>-haloalkyl,  
C<sub>1-6</sub>-alkoxy,  
C<sub>1-2</sub>-alkylamino,  
aminosulfonyl,  
C<sub>3-6</sub>-cycloalkyl,  
cyano,  
C<sub>1-2</sub>-hydroxyalkyl,  
nitro,  
C<sub>2-3</sub>-alkenyl,  
C<sub>2-3</sub>-alkynyl,  
C<sub>1-6</sub>-haloalkoxy,  
C<sub>1-6</sub>-carboxyalkyl,  
5-6-membered heterocyclyl-C<sub>1-6</sub>-alkylamino,  
~~unsubstituted or substituted phenyl and~~  
unsubstituted or substituted 5-6 membered heterocyclyl;

wherein R<sup>4</sup> is selected from a direct bond, C<sub>1-4</sub>-alkyl, and



wherein R<sup>2</sup> is selected from C<sub>1-2</sub>-alkyl, C<sub>2-6</sub>-branched alkyl, C<sub>2-4</sub>-branched haloalkyl, amino-C<sub>1-4</sub>-alkyl and C<sub>1-2</sub>-alkylamino-C<sub>1-2</sub>-alkyl;

wherein R<sup>e</sup> and R<sup>f</sup> are independently selected from H and C<sub>1-2</sub>-haloalkyl; and

wherein R<sup>7</sup> is selected from H, C<sub>1-3</sub>-alkyl, optionally substituted phenyl, optionally substituted phenyl-C<sub>1-3</sub>-alkyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted 4-6 membered heterocyclyl-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkoxy-C<sub>1-2</sub>-alkyl and C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkyl;

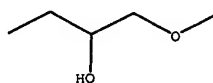
and pharmaceutically acceptable isomers and derivatives thereof.

Claim 46 (currently amended): A compound of Claim 45 wherein R is selected from 4-pyridyl, 3-pyridyl, 2-pyridyl, pyrimidinyl, triazolyl, pyridazinyl, indolyl, isoindolyl, indazolyl,

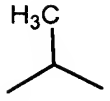
quinolyl, isoquinolyl, benzotriazolyl, naphthyridinyl and quinoxalyl, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, dimethylaminopropynyl, 1-methylpiperidinylmethoxy, dimethylaminoethoxyethoxy, methoxy and ethoxy; wherein R<sup>1</sup> is selected from 1,2-dihydroquinolyl, 1,2,3,4-tetrahydro-isoquinolyl, 2,3-dihydro-1H-indolyl, dihydro-benzimidazolyl, benzo[d]isothiazolyl, 2,3,4,4a,9,9a-hexahydro-1H-3-aza-fluorenyl, 5,6,7-trihydro-1,2,4-triazolo[3,4-a]isoquinolyl, and tetrahydroquinolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from bromo, chloro, fluoro, iodo, nitro, amino, cyano, aminoethyl, Boc-aminoethyl, hydroxy, oxo, aminosulfonyl, 4-methylpiperazinylsulfonyl, cyclohexyl, phenyl, phenylmethyl, morpholinylmethyl, 1-methylpiperazin-4-ylmethyl, 1-methylpiperazin-4-ylpropyl, morpholinylpropyl, piperidin-1-ylmethyl, 1-methylpiperidin-4-ylmethyl, 2-methyl-2-(1-methylpiperidin-4-yl)ethyl, morpholinylethyl, 1-(4-morpholinyl)-2,2-dimethylpropyl, piperidin-4-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-1-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-4-ylmethyl, 1-Boc-piperidin-4-ylmethyl, piperidin-4-ylpropyl, 1-Boc-piperidin-4-ylpropyl, piperidin-1-ylpropyl, pyrrolidin-1-ylpropyl, pyrrolidin-2-ylpropyl, 1-Boc-pyrrolidin-2-ylpropyl, pyrrolidin-1-ylmethyl, pyrrolidin-2-ylmethyl, 1-Boc-pyrrolidin-2-ylmethyl, pyrrolidinylpropenyl, pyrrolidinylbutenyl, fluorosulfonyl, methylsulfonyl, methylcarbonyl, Boc, piperidin-1-ylmethylcarbonyl, 4-methylpiperazin-1-ylcarbonylethyl, methoxycarbonyl, aminomethylcarbonyl, dimethylaminomethylcarbonyl, 3-ethoxycarbonyl-2-methyl-fur-5-yl, 4-methylpiperazin-1-yl, 4-methyl-1-piperidyl, 1-Boc-4-piperidyl, piperidin-4-yl, 1-methylpiperidin-4-yl, 1-methyl-(1,2,3,6-tetrahydropyridyl), imidazolyl, morpholinyl, 4-trifluoromethyl-1-piperidinyl, hydroxybutyl, methyl, ethyl, propyl, isopropyl, butyl, tert-butyl, sec-butyl, trifluoromethyl, pentafluoroethyl, nonafluorobutyl, dimethylaminopropyl, 1,1-di(trifluoromethyl)-1-hydroxymethyl, 1,1-di(trifluoromethyl)-1-(piperidinylethoxy)methyl, 1,1-di(trifluoromethyl)-1-(methoxyethoxyethoxy)methyl, 1-hydroxyethyl, 2-hydroxyethyl, trifluoromethoxy, 1-aminoethyl, 2-aminoethyl, 1-(N-isopropylamino)ethyl, 2-(N-isopropylamino)ethyl, dimethylaminoethoxy, 4-chlorophenoxy, phenyloxy, azetidin-3-ylmethoxy, 1-Boc-azetidin-3-ylmethoxy, pyrrol-2-ylmethoxy, 1-Boc-pyrrol-2-ylmethoxy, pyrrol-1-ylmethoxy, 1-methyl-pyrrol-2-ylmethoxy, 1-isopropyl-pyrrol-2-ylmethoxy, 1-Boc-piperidin-4-ylmethoxy, piperidin-4-ylmethoxy, 1-methylpiperidin-4-yloxy, isopropoxy, methoxy and ethoxy; wherein R<sup>2</sup> is selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, oxo, dimethylamino, aminosulfonyl, cyclopropyl, cyano, hydroxymethyl, nitro, propenyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, morpholinylethylamino, propynyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected from

thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl;

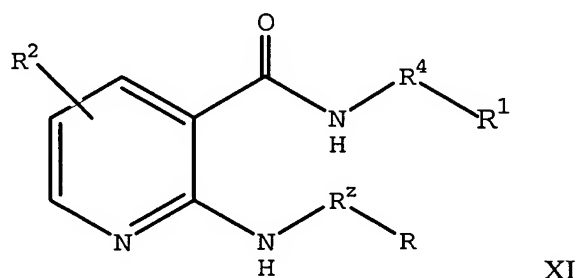
wherein R<sup>4</sup> is selected from a direct bond, ethyl, butyl, and



; and

wherein  $R^z$  is selected from methylenyl, ethylenyl, , and aminoethylenyl; and pharmaceutically acceptable derivatives thereof.

Claim 47 (currently amended): A compound of Claim ~~43~~ having Formula XI



wherein R is selected from

- a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and
- b) unsubstituted or substituted 9- or 10-membered fused heteroaryl,

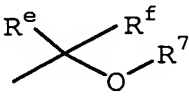
where substituted R is substituted with one or more substituents selected from halo, amino, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl,  $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkylamino, optionally substituted heterocyclyl- $C_{1-6}$ -alkyl,  $C_{1-6}$ -alkylamino- $C_{2-4}$ -alkynyl,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy- $C_{1-6}$ -alkoxy, and optionally substituted heterocyclyl- $C_{2-4}$ -alkynyl;

wherein  $R^1$  is selected from unsubstituted or substituted

- aryl,
- cycloalkyl,
- 5-6 membered heteroaryl and
- 9-10 membered bicyclic and 13-14 membered tricyclic heterocyclyl,

wherein substituted  $R^1$  is substituted with one or more substituents selected from halo,  $C_{1-6}$ -alkyl, optionally substituted  $C_{3-6}$ -cycloalkyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-4}$ -alkylenyl,  $C_{1-2}$ -haloalkoxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkyl, optionally substituted 4-6 membered heterocyclyl- $C_{2-4}$ -alkenyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted phenyloxy, optionally substituted 4-6 membered heterocyclyloxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkoxy, optionally substituted 4-6 membered heterocyclylsulfonyl, optionally substituted 4-6 membered heterocyclylamino, optionally substituted 4-6 membered heterocyclylcarbonyl, optionally substituted 5-6 membered heterocyclyl- $C_{1-4}$ -alkylcarbonyl,  $C_{1-2}$ -haloalkyl,  $C_{1-4}$ -aminoalkyl, nitro, amino, hydroxy, cyano, aminosulfonyl,  $C_{1-2}$ -

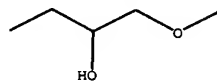
alkylsulfonyl, halosulfonyl, C<sub>1-4</sub>-alkylcarbonyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy, C<sub>1-4</sub>-alkoxycarbonyl, C<sub>1-4</sub>-

alkoxycarbonylamino-C<sub>1-4</sub>-alkyl, C<sub>1-4</sub>-hydroxyalkyl,  and C<sub>1-4</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

halo,  
hydroxy,  
amino,  
C<sub>1-6</sub>-alkyl,  
C<sub>1-6</sub>-haloalkyl,  
C<sub>1-6</sub>-alkoxy,  
C<sub>1-2</sub>-alkylamino,  
aminosulfonyl,  
C<sub>3-6</sub>-cycloalkyl,  
cyano,  
C<sub>1-2</sub>-hydroxyalkyl,  
nitro,  
C<sub>2-3</sub>-alkenyl,  
C<sub>2-3</sub>-alkynyl,  
C<sub>1-6</sub>-haloalkoxy,  
C<sub>1-6</sub>-carboxyalkyl,  
5-6-membered heterocyclyl-C<sub>1-6</sub>-alkylamino,  
~~unsubstituted or substituted phenyl~~ and  
unsubstituted or substituted 5-6 membered heterocyclyl;

wherein R<sup>4</sup> is selected from a direct bond, C<sub>1-4</sub>-alkyl, and



wherein R<sup>2</sup> is selected from C<sub>1-2</sub>-alkyl, C<sub>2-6</sub>-branched alkyl, C<sub>2-4</sub>-branched haloalkyl, amino-C<sub>1-4</sub>-alkyl and C<sub>1-2</sub>-alkylamino-C<sub>1-2</sub>-alkyl;

wherein R<sup>e</sup> and R<sup>f</sup> are independently selected from H and C<sub>1-2</sub>-haloalkyl; and

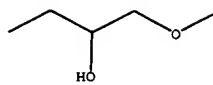
wherein R<sup>7</sup> is selected from H, C<sub>1-3</sub>-alkyl, optionally substituted phenyl, optionally substituted phenyl-C<sub>1-3</sub>-alkyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted 4-6 membered heterocyclyl-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkoxy-C<sub>1-2</sub>-alkyl and C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkyl;

and pharmaceutically acceptable isomers and derivatives thereof.

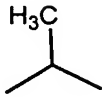
Claim 48 (currently amended): A compound of Claim 47 wherein R is selected from 4-pyridyl, 3-pyridyl, 2-pyridyl, pyrimidinyl, triazolyl, pyridazinyl, indolyl, isoindolyl, indazolyl,

quinolyl, isoquinolyl, benzotriazolyl, naphthyridinyl and quinozaliny, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, dimethylaminopropynyl, 1-methylpiperidinylmethoxy, dimethylaminoethoxyethoxy, methoxy and ethoxy; wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, indanyl, indenyl, naphthyl, cyclohexyl, isoxazolyl, pyrazolyl, thiazolyl, thiadiazolyl, thienyl, pyridyl, pyrimidinyl, pyridazinyl, 1,2-dihydroquinolyl, 1,2,3,4-tetrahydro-isoquinolyl, isoquinolyl, quinolyl, indolyl, isoindolyl, 2,3-dihydro-1H-indolyl, naphthyridinyl, quinozaliny, benzo[d]isothiazolyl, 2,3,4,4a,9,9a-hexahydro-1H-3-aza-fluorenyl, 5,6,7-trihydro-1,2,4-triazolo[3,4-a]isoquinolyl, tetrahydroquinolyl, indazolyl, 2,1,3-benzothiadiazolyl, benzodioxanyl, benzothienyl, benzofuryl, dihydro-benzimidazolyl, benzimidazolyl, benzoxazolyl and benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from bromo, chloro, fluoro, iodo, nitro, amino, cyano, aminoethyl, Boc-aminoethyl, hydroxy, oxo, aminosulfonyl, 4-methylpiperazinylsulfonyl, cyclohexyl, phenyl, phenylmethyl, morpholinylmethyl, 1-methylpiperazin-4-ylmethyl, 1-methylpiperazin-4-ylpropyl, morpholinylpropyl, piperidin-1-ylmethyl, 1-methylpiperidin-4-ylmethyl, 2-methyl-2-(1-methylpiperidin-4-yl)ethyl, morpholinylethyl, 1-(4-morpholinyl)-2,2-dimethylpropyl, piperidin-4-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-1-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-4-ylmethyl, 1-Boc-piperidin-4-ylmethyl, piperidin-4-ylpropyl, 1-Boc-piperidin-4-ylpropyl, piperidin-1-ylpropyl, pyrrolidin-1-ylpropyl, pyrrolidin-2-ylpropyl, 1-Boc-pyrrolidin-2-ylpropyl, pyrrolidin-1-ylmethyl, pyrrolidin-2-ylmethyl, 1-Boc-pyrrolidin-2-ylmethyl, pyrrolidinylpropenyl, pyrrolidinylbutenyl, fluorosulfonyl, methylsulfonyl, methylcarbonyl, Boc, piperidin-1-ylmethylcarbonyl, 4-methylpiperazin-1-ylcarbonylethyl, methoxycarbonyl, aminomethylcarbonyl, dimethylaminomethylcarbonyl, 3-ethoxycarbonyl-2-methyl-fur-5-yl, 4-methylpiperazin-1-yl, 4-methyl-1-piperidyl, 1-Boc-4-piperidyl, piperidin-4-yl, 1-methylpiperidin-4-yl, 1-methyl-(1,2,3,6-tetrahydropyridyl), imidazolyl, morpholinyl, 4-trifluoromethyl-1-piperidinyl, hydroxybutyl, methyl, ethyl, propyl, isopropyl, butyl, tert-butyl, sec-butyl, trifluoromethyl, pentafluoroethyl, nonafluorobutyl, dimethylaminopropyl, 1,1-di(trifluoromethyl)-1-hydroxymethyl, 1,1-di(trifluoromethyl)-1-(piperidinylethoxy)methyl, 1,1-di(trifluoromethyl)-1-(methoxyethoxyethoxy)methyl, 1-hydroxyethyl, 2-hydroxyethyl, trifluoromethoxy, 1-aminoethyl, 2-aminoethyl, 1-(N-isopropylamino)ethyl, 2-(N-isopropylamino)ethyl, dimethylaminoethoxy, 4-chlorophenoxy, phenyloxy, azetidin-3-ylmethoxy, 1-Boc-azetidin-3-ylmethoxy, pyrrol-2-ylmethoxy, 1-Boc-pyrrol-2-ylmethoxy, pyrrol-1-ylmethoxy, 1-methyl-pyrrol-2-ylmethoxy, 1-isopropyl-pyrrol-2-ylmethoxy, 1-Boc-piperidin-4-ylmethoxy, piperidin-4-ylmethoxy, 1-methylpiperidin-4-yloxy, isopropoxy, methoxy and ethoxy; wherein R<sup>2</sup> is selected from chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, oxo, dimethylamino, aminosulfonyl, cyclopropyl, cyano, hydroxymethyl, nitro, propenyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, morpholinylethylamino, propynyl, unsubstituted or substituted phenyl and unsubstituted or substituted heteroaryl selected from thienyl, furanyl, pyridyl, imidazolyl, and pyrazolyl;

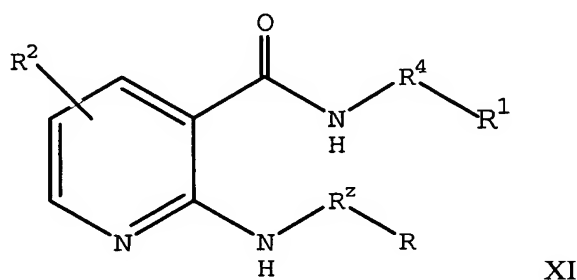
wherein  $R^4$  is selected from a direct bond, ethyl, butyl, and



; and

wherein  $R^z$  is selected from methylenyl, ethylenyl, , and aminoethylenyl; and pharmaceutically acceptable derivatives thereof.

Claim 49 (currently amended): A compound of Claim 43+ having Formula XI



wherein R is selected from

- a) unsubstituted or substituted 5- or 6-membered nitrogen-containing heteroaryl, and
- b) unsubstituted or substituted 9- or 10-membered fused heteroaryl,

where substituted R is substituted with one or more substituents selected from halo, amino, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl,  $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkylamino, optionally substituted heterocyclyl- $C_{1-6}$ -alkyl,  $C_{1-6}$ -alkylamino- $C_{2-4}$ -alkynyl,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy- $C_{1-6}$ -alkoxy, and optionally substituted heterocyclyl- $C_{2-4}$ -alkynyl;

wherein  $R^1$  is selected from unsubstituted or substituted

aryl,

cycloalkyl,

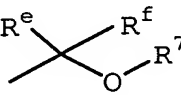
5-6 membered heteroaryl and

9-10 membered bicyclic and 13-14 membered tricyclic heterocyclyl,

wherein substituted  $R^1$  is substituted with one or more substituents selected from halo,  $C_{1-6}$ -alkyl,

optionally substituted  $C_{3-6}$ -cycloalkyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-4}$ -alkylenyl,  $C_{1-2}$ -haloalkoxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkyl, optionally substituted 4-6 membered heterocyclyl- $C_{2-4}$ -alkenyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted phenyloxy, optionally substituted 4-6 membered heterocyclyloxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkoxy, optionally substituted 4-6 membered heterocyclylsulfonyl, optionally substituted 4-6 membered heterocyclylamino, optionally substituted 4-6 membered

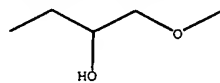
heterocyclylcarbonyl, optionally substituted 5-6 membered heterocyclyl-C<sub>1-4</sub>-alkylcarbonyl, C<sub>1-2</sub>-haloalkyl, C<sub>1-4</sub>-aminoalkyl, nitro, amino, hydroxy, cyano, aminosulfonyl, C<sub>1-2</sub>-alkylsulfonyl, halosulfonyl, C<sub>1-4</sub>-alkylcarbonyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy, C<sub>1-4</sub>-alkoxycarbonyl, C<sub>1-4</sub>-

alkoxycarbonylamino-C<sub>1-4</sub>-alkyl, C<sub>1-4</sub>-hydroxyalkyl,  and C<sub>1-4</sub>-alkoxy;

wherein R<sup>2</sup> is one or more substituents independently selected from

H,  
halo,  
hydroxy,  
amino,  
C<sub>1-6</sub>-alkyl,  
C<sub>1-6</sub>-haloalkyl,  
C<sub>1-6</sub>-alkoxy,  
C<sub>1-2</sub>-alkylamino,  
aminosulfonyl,  
C<sub>3-6</sub>-cycloalkyl,  
cyano,  
C<sub>1-2</sub>-hydroxyalkyl,  
nitro,  
C<sub>2-3</sub>-alkenyl,  
C<sub>2-3</sub>-alkynyl,  
C<sub>1-6</sub>-haloalkoxy,  
C<sub>1-6</sub>-carboxyalkyl,  
5-6-membered heterocyclyl-C<sub>1-6</sub>-alkylamino,  
~~unsubstituted or substituted phenyl~~ and  
unsubstituted or substituted 5-6 membered heterocyclyl;

wherein R<sup>4</sup> is selected from a direct bond, C<sub>1-4</sub>-alkyl, and



wherein R<sup>2</sup> is selected from C<sub>1-2</sub>-alkyl, C<sub>2-6</sub>-branched alkyl, C<sub>2-4</sub>-branched haloalkyl, amino-C<sub>1-4</sub>-alkyl and C<sub>1-2</sub>-alkylamino-C<sub>1-2</sub>-alkyl;

wherein R<sup>e</sup> and R<sup>f</sup> are independently selected from H and C<sub>1-2</sub>-haloalkyl; and

wherein R<sup>7</sup> is selected from H, C<sub>1-3</sub>-alkyl, optionally substituted phenyl, optionally substituted phenyl-C<sub>1-3</sub>-alkyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted 4-6 membered heterocyclyl-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkoxy-C<sub>1-2</sub>-alkyl and C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkyl;

provided R<sup>1</sup> is substituted with optionally substituted phenyloxy, optionally substituted 4-6 membered heterocyclyloxy, optionally substituted 4-6 membered heterocyclyl-C<sub>1-4</sub>-alkoxy, optionally substituted 4-6 membered heterocyclylsulfonyl, optionally substituted 4-6 membered heterocyclylamino, optionally substituted 4-6 membered heterocyclylcarbonyl, optionally substituted 4-6 membered heterocyclyl-C<sub>1-4</sub>-alkylcarbonyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy, or C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy; further provided R is not 3-pyridyl when R<sup>5</sup> R<sup>z</sup> is CH<sub>2</sub>; and pharmaceutically acceptable isomers and derivatives thereof.

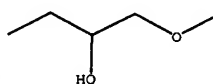
Claim 50 (currently amended): A compound of Claim 49 wherein R is selected from 4-pyridyl, 3-pyridyl, 2-pyridyl, pyrimidinyl, triazolyl, pyridazinyl, indolyl, isoindolyl, indazolyl, quinolyl, isoquinolyl, benzotriazolyl, naphthyridinyl and quinoxalinyl, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, dimethylaminopropynyl, 1-methylpiperdinylmethoxy, dimethylaminoethoxyethoxy, methoxy and ethoxy; wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, indanyl, indenyl, naphthyl, cyclohexyl, isoxazolyl, pyrazolyl, thiazolyl, thiadiazolyl, thienyl, pyridyl, pyrimidinyl, pyridazinyl, 1,2-dihydroquinolyl, 1,2,3,4-tetrahydroisoquinolyl, isoquinolyl, quinolyl, indolyl, isoindolyl, 2,3-dihydro-1H-indolyl, naphthyridinyl, quinoxalinyl, benzo[d]isothiazolyl, 2,3,4,4a,9,9a-hexahydro-1H-3-aza-fluorenyl, 5,6,7-trihydro-1,2,4-triazolo[3,4-a]isoquinolyl, tetrahydroquinolyl, indazolyl, 2,1,3-benzothiadiaazolyl, benzodioxanyl, benzothienyl, benzofuryl, benzimidazolyl, benzoxazolyl and benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from bromo, chloro, fluoro, iodo, nitro, amino, cyano, aminoethyl, Boc-aminoethyl, hydroxy, oxo, aminosulfonyl, 4-methylpiperazinylsulfonyl, cyclohexyl, phenyl, phenylmethyl, morpholinylmethyl, 1-methylpiperazin-4-ylmethyl, 1-methylpiperazin-4-ylpropyl, morpholinylpropyl, piperidin-1-ylmethyl, 1-methylpiperidin-4-ylmethyl, 2-methyl-2-(1-methylpiperidin-4-yl)ethyl, morpholinylethyl, 1-(4-morpholinyl)-2,2-dimethylpropyl, piperidin-4-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-1-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-4-ylmethyl, 1-Boc-piperidin-4-ylmethyl, piperidin-4-ylpropyl, 1-Boc-piperidin-4-ylpropyl, piperidin-1-ylpropyl, pyrrolidin-1-ylpropyl, pyrrolidin-2-ylpropyl, 1-Boc-pyrrolidin-2-ylpropyl, pyrrolidin-1-ylmethyl, pyrrolidin-2-ylmethyl, 1-Boc-pyrrolidin-2-ylmethyl, pyrrolidinylpropenyl, pyrrolidinylbutenyl, fluorosulfonyl, methylsulfonyl, methylcarbonyl, Boc, piperidin-1-ylmethylcarbonyl, 4-methylpiperazin-1-ylcarbonylethyl, methoxycarbonyl, aminomethylcarbonyl, dimethylaminomethylcarbonyl, 3-ethoxycarbonyl-2-methyl-fur-5-yl, 4-methylpiperazin-1-yl, 4-methyl-1-piperidyl, 1-Boc-4-piperidyl, piperidin-4-yl, 1-methylpiperidin-4-yl, 1-methyl-(1,2,3,6-tetrahydropyridyl), imidazolyl, morpholinyl, 4-trifluoromethyl-1-piperidinyl, hydroxybutyl, methyl, ethyl, propyl, isopropyl, butyl, tert-butyl, sec-butyl, trifluoromethyl, pentafluoroethyl, nonafluorobutyl, dimethylaminopropyl, 1,1-di(trifluoromethyl)-1-hydroxymethyl, 1,1-di(trifluoromethyl)-1-(piperidinylethoxy)methyl, 1,1-di(trifluoromethyl)-1-



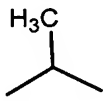
(methoxyethoxyethoxy)methyl, 1-hydroxyethyl, 2-hydroxyethyl, trifluoromethoxy, 1-aminoethyl, 2-aminoethyl, 1-(N-isopropylamino)ethyl, 2-(N-isopropylamino)ethyl, dimethylaminoethoxy, 4-chlorophenoxy, phenoxy, azetidin-3-ylmethoxy, 1-Boc-azetidin-3-ylmethoxy, pyrrol-2-ylmethoxy, 1-Boc-pyrrol-2-ylmethoxy, pyrrol-1-ylmethoxy, 1-methyl-pyrrol-2-ylmethoxy, 1-isopropyl-pyrrol-2-ylmethoxy, 1-Boc-piperidin-4-ylmethoxy, piperidin-4-ylmethoxy, 1-methylpiperidin-4-yloxy, isopropoxy, methoxy and ethoxy; wherein  $R^2$  is selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, oxo, dimethylamino, aminosulfonyl, cyclopropyl, cyano, hydroxymethyl, nitro, propenyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, morpholinylethylamino, propynyl, ~~unsubstituted or substituted phenyl~~ and unsubstituted or substituted heteroaryl selected from thienyl,

furanyl, pyridyl, imidazolyl, and pyrazolyl;

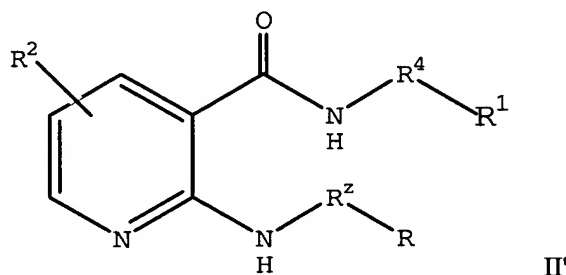
wherein  $R^4$  is selected from a direct bond, ethyl, butyl, and



; and

wherein  $R^z$  is selected from methylenyl, ethylenyl, , and aminoethylenyl; and pharmaceutically acceptable derivatives thereof.

Claim 51 (currently amended): A compound of Claim ~~431~~ having Formula II'



II'

wherein R is selected from

- unsubstituted or substituted 5- or 6-membered non-nitrogen-containing heterocyclyl, and
  - unsubstituted or substituted 9- or 10-membered fused partially unsaturated heterocyclyl,
- where R is substituted with one or more substituents selected from halo, amino, hydroxy,  $C_{1-6}$ -alkyl,  $C_{1-6}$ -haloalkyl,  $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkoxy, optionally substituted heterocyclyl- $C_{1-6}$ -alkylamino, optionally substituted heterocyclyl- $C_{1-6}$ -alkyl,  $C_{1-6}$ -alkylamino- $C_{2-4}$ -alkynyl,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy,  $C_{1-6}$ -alkylamino- $C_{1-6}$ -alkoxy- $C_{1-6}$ -alkoxy, and optionally substituted heterocyclyl- $C_{2-4}$ -alkynyl;

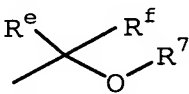
wherein  $R^1$  is selected from unsubstituted or substituted aryl,

cycloalkyl,

5-6 membered heteroaryl and

9-10 membered bicyclic and 13-14 membered tricyclic heterocyclyl,

wherein substituted  $R^1$  is substituted with one or more substituents selected from halo,  $C_{1-6}$ -alkyl, optionally substituted  $C_{3-6}$ -cycloalkyl, optionally substituted phenyl, optionally substituted phenyl- $C_{1-4}$ -alkylenyl,  $C_{1-2}$ -haloalkoxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkyl, optionally substituted 4-6 membered heterocyclyl- $C_{2-4}$ -alkenyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted phenyloxy, optionally substituted 4-6 membered heterocycliloxy, optionally substituted 4-6 membered heterocyclyl- $C_{1-4}$ -alkoxy, optionally substituted 4-6 membered heterocyclylsulfonyl, optionally substituted 4-6 membered heterocyclylamino, optionally substituted 4-6 membered heterocyclylcarbonyl, optionally substituted 5-6 membered heterocyclyl- $C_{1-4}$ -alkylcarbonyl,  $C_{1-2}$ -haloalkyl,  $C_{1-4}$ -aminoalkyl, nitro, amino, hydroxy, oxo,  $-NHC(O)NH_2$ , alkylcarbonylamino, cyano, aminosulfonyl,  $C_{1-2}$ -alkylsulfonyl, halosulfonyl,  $C_{1-4}$ -alkylcarbonyl,  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkyl,  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkoxy,  $C_{1-3}$ -alkylamino- $C_{1-3}$ -alkoxy- $C_{1-3}$ -alkoxy,  $C_{1-4}$ -alkoxycarbonyl,  $C_{1-4}$ -alkoxycarbonylamino- $C_{1-4}$ -alkyl,  $C_{1-4}$ -

hydroxyalkyl,  and  $C_{1-4}$ -alkoxy;

wherein  $R^2$  is one or more substituents independently selected from

H,

halo,

hydroxy,

amino,

$C_{1-6}$ -alkyl,

$C_{1-6}$ -haloalkyl,

$C_{1-6}$ -alkoxy,

$C_{1-2}$ -alkylamino,

aminosulfonyl,

$C_{3-6}$ -cycloalkyl,

cyano,

$C_{1-2}$ -hydroxyalkyl,

nitro,

$C_{2-3}$ -alkenyl,

$C_{2-3}$ -alkynyl,

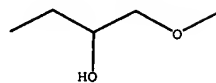
$C_{1-6}$ -haloalkoxy,

$C_{1-6}$ -carboxyalkyl,

5-6-membered heterocyclyl- $C_{1-6}$ -alkylamino,

~~unsubstituted or substituted phenyl~~ and

unsubstituted or substituted 5-6 membered heterocyclyl;



wherein R<sup>4</sup> is selected from a direct bond, C<sub>1-4</sub>-alkyl, and

wherein R<sup>z</sup> is selected from C<sub>1-2</sub>-alkyl, C<sub>2-6</sub>-branched alkyl, C<sub>2-4</sub>-branched haloalkyl, amino-C<sub>1-4</sub>-alkyl and C<sub>1-2</sub>-alkylamino-C<sub>1-2</sub>-alkyl;

wherein R<sup>e</sup> and R<sup>f</sup> are independently selected from H and C<sub>1-2</sub>-haloalkyl; and

wherein R<sup>7</sup> is selected from H, C<sub>1-3</sub>-alkyl, optionally substituted phenyl-C<sub>1-3</sub>-alkyl, optionally substituted 4-6 membered heterocyclyl, optionally substituted 4-6 membered heterocyclyl-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkylamino-C<sub>1-3</sub>-alkyl, C<sub>1-3</sub>-alkoxy-C<sub>1-2</sub>-alkyl and C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkoxy-C<sub>1-3</sub>-alkyl;

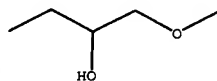
and pharmaceutically acceptable isomers and derivatives thereof.

Claim 52 (currently amended): A compound of Claim 50 wherein R is selected from 2,3-dihydrobenzofuryl, and tetrahydropyran, where R is unsubstituted or substituted with one or more substituents selected from chloro, fluoro, amino, hydroxy, methyl, ethyl, propyl, trifluoromethyl, dimethylaminopropynyl, 1-methylpiperdinylmethoxy, dimethylaminoethoxyethoxy, methoxy and ethoxy; wherein R<sup>1</sup> is selected from phenyl, tetrahydronaphthyl, indanyl, indenyl, naphthyl, cyclohexyl, isoxazolyl, pyrazolyl, thiazolyl, thiadiazolyl, thienyl, pyridyl, pyrimidinyl, pyridazinyl, 1,2-dihydroquinolyl, 1,2,3,4-tetrahydro-isoquinolyl, isoquinolyl, quinolyl, indolyl, isoindolyl, 2,3-dihydro-1H-indolyl, naphthyridinyl, quinoxalinyl, benzo[d]isothiazolyl, 2,3,4,4a,9,9a-hexahydro-1H-3-aza-fluorenyl, 5,6,7-trihydro-1,2,4-triazolo[3,4-a]isoquinolyl, tetrahydroquinolyl, indazolyl, 2,1,3-benzothiadiazolyl, benzodioxanyl, benzothienyl, benzofuryl, benzimidazolyl, dihydro-benzimidazolyl, benzoxazolyl and benzthiazolyl, where R<sup>1</sup> is unsubstituted or substituted with one or more substituents selected from bromo, chloro, fluoro, iodo, nitro, amino, cyano, aminoethyl, Boc-aminoethyl, hydroxy, oxo, aminosulfonyl, 4-methylpiperazinylsulfonyl, cyclohexyl, phenyl, phenylmethyl, morpholinylmethyl, 1-methylpiperazin-4-ylmethyl, 1-methylpiperazin-4-ylpropyl, morpholinylpropyl, piperidin-1-ylmethyl, 1-methylpiperidin-4-ylmethyl, 2-methyl-2-(1-methylpiperidin-4-yl)ethyl, morpholinylethyl, 1-(4-morpholinyl)-2,2-dimethylpropyl, piperidin-4-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-1-ylethyl, 1-Boc-piperidin-4-ylethyl, piperidin-4-ylmethyl, 1-Boc-piperidin-4-ylmethyl, piperidin-4-ylpropyl, 1-Boc-piperidin-4-ylpropyl, piperidin-1-ylpropyl, pyrrolidin-1-ylpropyl, pyrrolidin-2-ylpropyl, 1-Boc-pyrrolidin-2-ylpropyl, pyrrolidin-1-ylmethyl, pyrrolidin-2-ylmethyl, 1-Boc-pyrrolidin-2-ylmethyl, pyrrolidinylpropenyl, pyrrolidinylbutenyl, fluorosulfonyl, methylsulfonyl, methylcarbonyl, Boc, piperidin-1-ylmethylcarbonyl, 4-methylpiperazin-1-ylcarbonylethyl, methoxycarbonyl, aminomethylcarbonyl, dimethylaminomethylcarbonyl, 3-ethoxycarbonyl-2-methyl-fur-5-yl, 4-methylpiperazin-1-yl, 4-methyl-1-piperidyl, 1-Boc-4-piperidyl, piperidin-4-yl, 1-methylpiperidin-4-yl, 1-methyl-(1,2,3,6-tetrahydropyridyl), imidazolyl, morpholinyl, 4-trifluoromethyl-1-piperidinyl, hydroxybutyl, methyl, ethyl, propyl, isopropyl, butyl, tert-butyl, sec-butyl, trifluoromethyl, pentafluoroethyl,

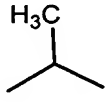
nonafluorobutyl, dimethylaminopropyl, 1,1-di(trifluoromethyl)-1-hydroxymethyl, 1,1-di(trifluoromethyl)-1-(piperidinylethoxy)methyl, 1,1-di(trifluoromethyl)-1-(methoxyethoxyethoxy)methyl, 1-hydroxyethyl, 2-hydroxyethyl, trifluoromethoxy, 1-aminoethyl, 2-aminoethyl, 1-(N-isopropylamino)ethyl, 2-(N-isopropylamino)ethyl, dimethylaminoethoxy, 4-chlorophenoxy, phenyloxy, azetidin-3-ylmethoxy, 1-Boc-azetidin-3-ylmethoxy, pyrrol-2-ylmethoxy, 1-Boc-pyrrol-2-ylmethoxy, pyrrol-1-ylmethoxy, 1-methyl-pyrrol-2-ylmethoxy, 1-isopropyl-pyrrol-2-ylmethoxy, 1-Boc-piperdin-4-ylmethoxy, piperdin-4-ylmethoxy, 1-methylpiperdin-4-yloxy, isopropoxy, methoxy and ethoxy; wherein  $R^2$  is selected from H, chloro, fluoro, bromo, amino, hydroxy, methyl, ethyl, propyl, oxo, dimethylamino, aminosulfonyl, cyclopropyl, cyano, hydroxymethyl, nitro, propenyl, trifluoromethyl, methoxy, ethoxy, trifluoromethoxy, carboxymethyl, morpholinylethylamino, propynyl, ~~unsubstituted or substituted phenyl~~ and unsubstituted or substituted heteroaryl selected from thienyl,

furanyl, pyridyl, imidazolyl, and pyrazolyl;

wherein  $R^4$  is selected from a direct bond, ethyl, butyl, and



; and

wherein  $R^z$  is selected from methylenyl, ethylenyl, , and aminoethylenyl; and pharmaceutically acceptable derivatives thereof.

Claim 53 (canceled).

Claim 54 (canceled).

Claim 55 (canceled).

Claim 56 (withdrawn): Compound of Claim 1 wherein ring A is selected from dihydropyran, dihydrothienyl, dihydrofuryl, oxo-dihydrofuryl, pyrrolinyl, dihydrothiazolyl, dihydro-oxazolyl, dihydro-isothiazolyl, dihydro-isoxazolyl, imidazolynyl, pyrazolynyl, triazinyl, thienyl, furanyl, pyrrolyl, thiazolyl, oxazolyl, imidazolyl, pyrazolyl, isoxazolyl, triazolyl and isothiazolyl.

Claim 57 (withdrawn): Compound of Claim 1 wherein R is selected from substituted or unsubstituted, saturated or partially saturated 5-6 membered heterocyclyl, and substituted or unsubstituted saturated or partially saturated fused 9-, 10- or 11-membered heterocyclyl.

Claim 58 (canceled).

Claim 59 (canceled).

Claim 60 (canceled).

Claim 61 (canceled).

Claim 62 (canceled).

Claim 63 (currently amended): A pharmaceutical composition comprising a pharmaceutically-acceptable inert carrier and an effective amount of a compound from any one of Claims 43-~~62-55~~ and 70-84.

Claim 64 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Claim 43.

Claim 65 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Claim 45.

Claim 66 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Claim 47.

Claim 67 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Claim 49.

Claim 68 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Claim 51.

Claim 69 (withdrawn): A method of treating cancer in a subject, said method comprising administering an effective amount of a compound of Claim 53.

Claim 70 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-(3-(4-piperidinyloxy)-5-(trifluoromethyl)phenyl)-2-((2-(3-pyridinyl)ethyl)amino)-3-pyridinecarboxamide.

Claim 71 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is {6-fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}-N-[4-(isopropyl)phenyl]carboxamide.

Claim 72 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-[3-(pyrrolidin-2-yl-methoxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-yl-methyl)-amino]-nicotinamide.

Claim 73 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-(1-acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-[(2,3-dihydro-benzofuran-5-ylmethyl)-amino]-nicotinamide.

Claim 74 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-(3,3-dimethyl-2,3-dihydro-benzofuran-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide.

Claim 75 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-[3-(1-methyl-pyrrolidin-2-ylmethoxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide.

Claim 76 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is 2-[(pyridin-4-ylmethyl)-amino]-N-[3-(2-pyrrolidin-1-yl-ethoxy)-4-trifluoromethyl-phenyl]-nicotinamide.

Claim 77 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-(3,3-dimethylindolin-6-yl){2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide.

Claim 78 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-[3-(pyrrolidin-2-yl-methoxy)-4-pentafluoroethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide.

Claim 79 (new): Compound of Claim 43 and pharmaceutically acceptable salts thereof wherein said compound is N-[3-(azetidin-3-ylmethoxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide.

Claim 80 (new): Compound of Claim 47 and pharmaceutically acceptable derivatives thereof selected from

5-Bromo-N-[2-(4-chlorophenyl)ethyl]-2-[(pyridin-4-ylmethyl)amino](3-pyridyl)carboxamide;  
 {6-Chloro-5-fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}-N-[4-(isopropyl)phenyl]carboxamide;  
 {5-Fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}-N-[4-(isopropyl)phenyl]carboxamide;

N-(3,4-Dichlorophenyl){6-[(2-morpholin-4-ylethyl)amino]-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-[4-(*tert*-Butyl)phenyl]{6-fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 {6-Fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}-N-[4-(isopropyl)phenyl]carboxamide;  
 {6-Fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}-N-[3-(trifluoromethyl)phenyl]carboxamide;  
 N-(1-Bromo-(3-isoquinolyl)){6-fluoro-2-[(4-pyridylmethyl)amino](3-pyridyl)}-carboxamide;  
 N-(4-chlorophenyl){6-methyl-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-(3,4-dichlorophenyl){6-methyl-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-(3-fluoro-4-methylphenyl){6-methyl-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-(3,4-dichlorophenyl){6-chloro-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-(4-chlorophenyl){6-chloro-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-(3-chlorophenyl){6-chloro-2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-(4-chlorophenyl){2-[(4-pyridylethyl)amino]-5-(3-thienyl)-(3-pyridyl)}carboxamide; and  
 N-(4-chlorophenyl){5-(4-methoxyphenyl)-2-[(4-pyridylmethyl)amino]-(3-pyridyl)}carboxamide.

Claim 81 (new): Compound of Claim 49 and pharmaceutically acceptable derivatives thereof selected from

N-(4-Phenoxyphenyl){2-[(2-(3-pyridyl)ethyl)amino](3-pyridyl)}carboxamide;  
 N-(4-Phenoxyphenyl){2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide hydrochloride;  
 N-(3-Phenoxyphenyl){2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide hydrochloride;  
 N-[2-(4-Phenoxyphenyl)ethyl]-2-[(pyridin-4-ylmethyl)amino](3-pyridyl)carboxamide;  
 N-(2-Hydroxy-3-phenoxypropyl)-2-[(pyridin-4-ylmethyl)amino](3-pyridyl)carboxamide;  
 N-{3-[(4-Methylpiperaziny)sulfonyl]phenyl}{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 N-{2-[2-(dimethylamino)ethoxy]-5-(*tert*-butyl)phenyl}{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;  
 {2-[(4-Pyridylmethyl)amino](3-pyridyl)}-N-{4-[2,2,2-trifluoro-1-(2-piperidylethoxy)-1-(trifluoromethyl)ethyl]phenyl}carboxamide;  
 N-[3-(4-Boc-piperazine-1-carbonyl)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-[3-(4-Boc-piperazine-1-carbonyl)-5-trifluoromethyl-phenyl]-2-(2-pyridin-4-yl-ethylamino)-nicotinamide;  
 2-[(2-Methoxy-pyridin-4-ylmethyl)-amino]-N-[3-(1-Boc-piperidin-4-ylmethoxy)-5-trifluoromethyl-phenyl]-nicotinamide;  
 2-[(2-Methoxy-pyridin-4-ylmethyl)-amino]-N-[3-(1-Boc-piperidin-4-ylmethoxy)-5-trifluoromethyl-phenyl]-nicotinamide;  
 2-[(2-Methoxy-pyridin-4-ylmethyl)-amino]-N-[3-(1-methyl-pyrrolidin-2-ylmethoxy)-5-trifluoromethyl-phenyl]-nicotinamide;

(R) N-[3-(1-Boc-pyrrolidin-2-ylmethoxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

(R) N-[3-(1-Methyl-pyrrolidin-2-ylmethoxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3-(1-Methyl-piperidin-4-yloxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3-tert-Butyl-4-(1-Boc-pyrrolidin-2-ylmethoxy)-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3-(1-Boc-azetidin-3-ylmethoxy)-5-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

2-[(2-Methoxy-pyridin-4-ylmethyl)-amino]-N-[3-(1-Boc-azetidin-3-ylmethoxy)-5-trifluoromethyl-phenyl]-nicotinamide;

(R) N-[3-(2-Hydroxy-3-pyrrolidin-1-yl-propoxy)-4-pentafluoroethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

(S) N-[3-(2-Hydroxy-3-pyrrolidin-1-yl-propoxy)-4-pentafluoroethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[4-tert-Butyl-3-(1-methyl-piperidin-4-ylmethoxy)-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3-(1-Methyl-piperidin-4-ylmethoxy)-4-pentafluoroethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[4-Pentafluoroethyl-3-(2-piperidin-1-yl-ethoxy)-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[4-Trifluoromethyl-3-(2-piperidin-1-yl-ethoxy)-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

(S) N-[3-(1-Boc-pyrrolidin-2-ylmethoxy)-4-pentafluoroethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

(R) N-[3-(1-Boc-pyrrolidin-2-ylmethoxy)-4-trifluoromethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3,3-Dimethyl-1-(pyrrolidin-2-ylmethoxy)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

(R) N-[3-(1-Boc-pyrrolidin-2-ylmethoxy)-4-pentafluoroethyl-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[4-tert-Butyl-3-(1-methyl-azetidin-3-ylmethoxy)-phenyl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

2-[(2-Methoxy-pyridin-4-ylmethyl)-amino]-N-[3-(1-Boc-piperidin-4-yloxy)-5-trifluoromethyl-phenyl]-nicotinamide; and

2-[(2-Methoxy-pyridin-4-ylmethyl)-amino]-N-[3-(piperidin-4-ylmethoxy)-5-trifluoromethyl-phenyl]-nicotinamide.



Claim 82 (new): Compound of Claim 45 and pharmaceutically acceptable derivatives thereof selected from

2-[(Pyridin-4-ylmethyl)-amino]-N-(2,2,4-trimethyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-nicotinamide;

N-(4-Acetyl-2,2-dimethyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-(2,2-Dimethyl-3-oxo-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-(4,4-Dimethyl-1-oxo-1,2,3,4-tetrahydro-isoquinolin-7-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-(1-Acetylinolin-6-yl){2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-Indolin-6-yl{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-[1-(1-Methyl-(4-piperidyl))indolin-6-yl]{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-[1-(1-Methyl-(4-piperidyl))indolin-6-yl]{2-[(2-(3-pyridyl)ethyl)amino](3-pyridyl)}carboxamide;

N-[1-(2-Piperidylethyl)indolin-6-yl]{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-[1-(2-Piperidylacetyl)indolin-6-yl]{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-[3,3-Dimethyl-1-(1-methyl(4-piperidyl))indolin-6-yl]{2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-(3,3-Dimethylindolin-6-yl){2-[(4-pyridylmethyl)amino](3-pyridyl)}carboxamide;

N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3,3-Dimethyl-1-(1-methyl-piperidin-4-yl)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-(1-Boc-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3,3-Dimethyl-1-(1-Boc-piperidin-4-ylmethyl)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3,3-Dimethyl-1-(1-methyl-piperidin-4-yl)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[1-(2-Dimethylamino-acetyl)-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[1-(2-Dimethylamino-acetyl)-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3,3-Dimethyl-1-(1-Boc-pyrrolidin-2-ylmethoxy)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;

N-[3,3-Dimethyl-1-(2-Boc-amino-acetyl)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-[3,3-Dimethyl-1-(2-Boc-amino-acetyl)-2,3-dihydro-1H-indol-6-yl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-[[2-(3-morpholin-4-yl-propylamino)-pyridin-4-ylmethyl]-amino]-nicotinamide;  
 2-[(Pyridin-4-ylmethyl)-amino]-N-(3,9,9-trimethyl-2,3,4,4a,9,9a-hexahydro-1H-3-aza-fluoren-6-yl)-nicotinamide;  
 N-[3,3-Dimethyl-1-(1-Boc-piperidin-4-ylmethyl)-2,3-dihydro-1H-indol-6-yl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-[3,3-Dimethyl-1-(1-methyl-piperidin-4-ylmethyl)-2,3-dihydro-1H-indol-6-yl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(3,3-Dimethyl-1,1-dioxo-2,3-dihydro-1H-1 $\lambda$ -benzo[d]isothiazol-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(3,3-Dimethyl-2,3-dihydro-benzofuran-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(3,3-Dimethyl-2,3-dihydro-1H-indol-6-yl)-2-({2-[2-(1-methyl-piperidin-4-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-nicotinamide;  
 N-(2,2-Dimethyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(4,4-Dimethyl-1,2,3,4-tetrahydro-isoquinolin-7-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(3,3-Dimethyl-1-piperidin-4-yl-2,3-dihydro-1H-indol-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-({2-[2-(1-methyl-pyrrolidin-2-yl)-ethylamino]-pyrimidin-4-ylmethyl}-amino)-nicotinamide;  
 N-(3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-[3,3-Dimethyl-1-(piperidin-4-ylmethyl)-2,3-dihydro-1H-indol-6-yl]-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(3,3-Dimethyl-1-piperidin-4-yl-2,3-dihydro-1H-indol-6-yl)-2-[(2-methoxy-pyridin-4-ylmethyl)-amino]-nicotinamide;  
 N-(2,3,3-Trimethyl-1,1-dioxo-2,3-dihydro-1H-1 $\lambda$ -benzo[d]isothiazol-6-yl)-2-[(pyridin-4-ylmethyl)-amino]-benzamide;  
 N-[3,3-Dimethyl-1,1-dioxo-2-(2-piperidin-1-yl-ethyl)-2,3-dihydro-1H-1 $\lambda$ '-benzo[d]isothiazol-6-yl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide; and  
 N-[2-(2-Dimethylamino-ethyl)-3,3-dimethyl-1,1-dioxo-2,3-dihydro-1H-1 $\lambda$ '-benzo[d]isothiazol-6-yl]-2-[(pyridin-4-ylmethyl)-amino]-nicotinamide.

Claim 83 (new): Compound of Claim 43 and pharmaceutically acceptable derivatives thereof selected from

N-(4-tert-Butyl-phenyl)-2-{{2-(1-methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-  
 nicotinamide;  
 N-(3-Trifluoromethyl-phenyl)-2-{{2-(1-methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-  
 nicotinamide;  
 N-(3-tert-Butyl-isoxazol-5-yl)-2-{{2-(1-methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-  
 nicotinamide;  
 N-[4-(tert-Butyl)phenyl]{2-[(2-[(1-methyl(4-piperidyl))-methoxy](4-pyridyl))methyl]amino}(3-  
 pyridyl)}carboxamide;  
 (2-[(2-[(2-[(Dimethylamino)ethoxy]ethoxy)(4-pyridyl))methyl]amino}(3-pyridyl))-N-[4-(tert-  
 butyl)phenyl]carboxamide;  
 (2-[(2-[(2-[(Dimethylamino)ethoxy]ethoxy)(4-pyridyl))methyl]amino}-6-fluoro(3-pyridyl))-N-[3-  
 (trifluoromethyl)phenyl]carboxamide;  
 2-{{2-(1-Isopropyl-azetidin-3-ylmethoxy)-pyridin-4-ylmethyl}-amino}-N-(4-trifluoromethyl-phenyl)-  
 nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-{{2-(1-isopropyl-azetidin-3-ylmethoxy)-pyridin-4-ylmethyl}-amino}-  
 nicotinamide;  
 N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-({2-[2-(1-methyl-piperidin-4-yl)-ethoxy]-  
 pyridin-4-ylmethyl}-amino)-nicotinamide;  
 2-({2-[2-(1-Methyl-piperidin-4-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-N-(3-trifluoromethyl-  
 phenyl)-nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-({2-[2-(1-methyl-pyrrolidin-2-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-  
 nicotinamide;  
 2-({2-[2-(1-Methyl-pyrrolidin-2-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-N-(4-pentafluoroethyl-  
 phenyl)-nicotinamide;  
 N-(4-Pentafluoroethyl-phenyl)-2-{{2-(2-pyrrolidin-1-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-  
 nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-{{2-(2-pyrrolidin-1-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 2-{{2-(1-Methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl}-amino}-N-(4-trifluoromethyl-phenyl)-  
 nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-{{2-(1-methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl}-amino}-  
 nicotinamide;  
 2-({2-[3-(1-Methyl-piperidin-4-yl)-propoxy]-pyridin-4-ylmethyl}-amino)-N-(4-pentafluoroethyl-  
 phenyl)-nicotinamide;  
 2-{{2-(3-Morpholin-4-yl-propoxy)-pyridin-4-ylmethyl}-amino}-N-(4-pentafluoroethyl-phenyl)-  
 nicotinamide;  
 (S) 2-{{2-(1-Methyl-pyrrolidin-2-ylmethoxy)-pyridin-4-ylmethyl}-amino}-N-(4-pentafluoroethyl-  
 phenyl)-nicotinamide;

N-(3-tert-Butyl-isoxazol-5-yl)-2-{{2-(3-morpholin-4-yl-propoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-{{2-(3-morpholin-4-yl-propoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-{{2-(2-morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 2-{{2-(2-Morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-N-(4-trifluoromethyl-phenyl)-nicotinamide;  
 2-{{2-(2-Morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-N-(3-trifluoromethyl-phenyl)-nicotinamide;  
 2-{{2-(2-Morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-N-(4-pentafluoroethyl-phenyl)-nicotinamide;  
 N-(3-tert-Butyl-isoxazol-5-yl)-2-{{2-(2-morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(2-morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(1-methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 2-{{2-(1-Methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-N-(4-trifluoromethyl-phenyl)-nicotinamide;  
 2-{{2-(1-Methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-N-(4-pentafluoroethyl-phenyl)-nicotinamide;  
 2-{{2-(1-Methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl}-amino}-N-(4-tert-butyl-phenyl)-nicotinamide;  
 (R) N-(4-tert-Butyl-phenyl)-2-{{2-(1-methyl-pyrrolidin-2-ylmethoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 N-(3,3-Dimethyl-2,3-dihydro-benzofuran-6-yl)-2-{{2-(1-methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 2-{{2-[3-(1-Methyl-piperidin-4-yl)-propoxy]-pyridin-4-ylmethyl}-amino}-N-(4-trifluoromethyl-phenyl)-nicotinamide;  
 2-{{2-[3-(1-Methyl-piperidin-4-yl)-propoxy]-pyridin-4-ylmethyl}-amino}-N-(3-trifluoromethyl-phenyl)-nicotinamide;  
 2-{{2-[3-(1-Methyl-piperidin-4-yl)-propoxy]-pyridin-4-ylmethyl}-amino}-N-(4-tert-butyl-phenyl)-nicotinamide;  
 2-{{2-[3-(1-Methyl-piperidin-4-yl)-propoxy]-pyridin-4-ylmethyl}-amino}-N-(3-tert-butyl-isoxazol-5-yl)-nicotinamide;  
 N-(3,3-Dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-[3-(1-methyl-piperidin-4-yl)-propoxy]-pyridin-4-ylmethyl}-amino}-nicotinamide;  
 2-{{2-(1-Methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl}-amino}-N-(4-pentafluoroethyl-phenyl)-nicotinamide;

N-(3-tert-Butyl-isoxazol-5-yl)-2-{{2-(1-methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl]-amino}-nicotinamide;  
 N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(1-methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl]-amino}-nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-{{2-(3-morpholin-4-yl-propylamino)-pyrimidin-4-ylmethyl]-amino}-nicotinamide;  
 2-{{2-(3-Morpholin-4-yl-propylamino)-pyrimidin-4-ylmethyl]-amino}-N-(4-pentafluoroethyl-phenyl)-nicotinamide;  
 2-{{2-(3-Morpholin-4-yl-propylamino)-pyrimidin-4-ylmethyl]-amino}-N-(3-trifluoromethyl-phenyl)-nicotinamide;  
 N-(4-tert-Butyl-phenyl)-2-({2-[2-(1-methyl-pyrrolidin-2-yl)-ethylamino]-pyrimidin-4-ylmethyl}-amino)-nicotinamide;  
 N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-({2-[2-(1-methyl-pyrrolidin-2-yl)-ethylamino]-pyrimidin-4-ylmethyl}-amino)-nicotinamide;  
 2-{{2-(1-Methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl]-amino}-N-[3-(1-methyl-piperidin-4-yl)-5-trifluoromethyl-phenyl]-nicotinamide;  
 N-(3-tert-Butyl-isoxazol-5-yl)-2-{{2-(1-methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl]-amino}-nicotinamide;  
 2-{{2-(1-Benzhydryl-azetidin-3-yloxy)-pyridin-4-ylmethyl]-amino}-N-(4-tert-butyl-phenyl)-nicotinamide.  
 N-(4-tert-Butyl-phenyl)-2-({2-[2-(1-methyl-piperidin-4-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-nicotinamide;  
 N-(3-tert-Butyl-isoxazol-5-yl)-2-({2-[2-(1-methyl-piperidin-4-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-nicotinamide;  
 N-(3-trifluoromethylphenyl)-2-({2-[2-(1-methyl-piperidin-4-yl)-ethoxy]-pyridin-4-ylmethyl}-amino)-nicotinamide;  
 N-(3,3-Dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(1-methyl-piperidin-4-ylmethoxy)-pyridin-4-ylmethyl]-amino}-nicotinamide;  
 N-(3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(2-morpholin-4-yl-ethoxy)-pyridin-4-ylmethyl]-amino}-nicotinamide;  
 N-(3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(1-methyl-piperidin-4-yloxy)-pyridin-4-ylmethyl]-amino}-nicotinamide;  
 N-(3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-{{2-(2-morpholin-4-yl-propoxy)-pyridin-4-ylmethyl]-amino}-nicotinamide; and  
 2-{{2-(azetidin-3-yloxy)-pyridin-4-ylmethyl]-amino}-N-(4-tert-butyl-phenyl)nicotinamide.

Claim 84 (new): Compound of Claim 51 and pharmaceutically acceptable derivatives thereof selected from

2-[(2,3-Dihydro-benzofuran-5-ylmethyl)-amino]-N-{4-[1-methyl-1-(1-methyl-piperidin-4-yl)-ethyl]-phenyl}-nicotinamide;  
N-(1-Acetyl-3,3-dimethyl-2,3-dihydro-1H-indol-6-yl)-2-[(2,3-dihydro-benzofuran-5-ylmethyl)-amino]-nicotinamide;  
2-[(2,3-Dihydro-benzofuran-5-ylmethyl)-amino]-N-[3,3-dimethyl-1-(1-Boc-piperidin-4-ylmethyl)-2,3-dihydro-1H-indol-6-yl]-nicotinamide;  
2-[(2,3-Dihydro-benzofuran-5-ylmethyl)-amino]-N-[3,3-dimethyl-1-(1-methylpiperidin-4-ylmethyl)-2,3-dihydro-1H-indol-6-yl]-nicotinamide; and  
2-[(2,3-Dihydro-benzofuran-6-ylmethyl)-amino]-N-[3-(1-Boc-pyrrolidin-2-ylmethoxy)-4-pentafluoroethyl-phenyl]-nicotinamide.